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(57) Abstract

As a step towards understanding the complex differences between normal and cancer cells, gene expression patterns were examined in gastrointestinal tumors. More than 300,000 transcripts derived from at least 45,000 different genes were analyzed. Although extensive similarity was noted between the expression profiles, more than 500 transcripts that were expressed at significantly different levels in normal and neoplastic cells were identified. These data provide insights into the extent of expression differences underlying malignancy and reveal genes that are useful as diagnostic or prognostic markers.

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Gene Expression Profiles in Normal and Cancer Cells

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TECHNICAL FIELD OF THE INVENTION

This invention is related to the diagnosis of cancer, and tools for carrying out such diagnosis.

BACKGROUND OF THE INVENTION

Much of cancer research over the past 50 years has been devoted to the analyses of genes that are expressed differently in tumor cells compared to their normal counterparts. Although hundreds of studies have pointed out differences in the expression of one or a few genes, no comprehensive study of gene expression in the cancer cell has been reported. It is therefore not known how many genes are expressed differentially in tumor versus normal cells, whether the bulk of these differences are cell autonomous rather than being dependent on the tumor microenvironment, and whether most differences are cell-type specific or tumor specific. Thus there is a need in the art for information on the molecular changes that occur in cells during cancer development and progression.

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SUMMARY OF THE INVENTION

According to one embodiment of the invention, a method is provided for diagnosing colon cancer in a sample suspected of being neoplastic. The method comprises the steps of:

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comparing the level of at least one transcript in a first sample of a tissue to a second sample, wherein the first sample is of a colonic tissue suspected of being neoplastic and the second sample is of a normal human colonic tissue, and wherein the transcript is identified by a tag selected from the group consisting of those shown in Table 3;

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identifying the first sample as neoplastic when the level of the at least one transcript is found to be lower in the first sample than in the second sample.

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According to another embodiment of the invention, another method is provided for diagnosing colon cancer in a sample suspected of being neoplastic. The method comprises the steps of:

comparing the level of at least one transcript in a first sample of a tissue to a second sample, wherein the first sample is of a colonic tissue suspected of being neoplastic and the second sample is of a normal human colonic tissue, and wherein the transcript is identified by a tag selected from the group consisting of those shown in Table 2;

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identifying the first sample as neoplastic when the level of the at least one transcript is found to be higher in the first sample than in the second sample.

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In another embodiment of the invention an isolated and purified human nucleic acid molecule is provided. The molecule comprises a SAGE tag selected from SEQ ID NO:1-732.

In yet another aspect of the invention an isolated nucleotide probe is provided. The probe comprises at least 12 nucleotides of a human nucleic acid molecule, wherein the human nucleic acid molecule comprises a SAGE tag selected from SEQ ID NO: 1-732.

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According to another aspect of the invention a method is provided for diagnosing pancreatic cancer in a sample suspected of being neoplastic. The method comprises the steps of:

comparing the level of at least one transcript in a first sample of a tissue to a second sample, wherein the first sample is of a pancreatic tissue suspected of being neoplastic and the second sample is of a normal human colon tissue, wherein said transcript is identified by a tag selected from the group consisting of those shown Table 4;

identifying the first sample as neoplastic when the level of the at least one transcript is found to be higher in the first sample than in the second sample.

According to still another embodiment of the invention a method of diagnosing cancer in a sample suspected of being neoplastic is provided. The method comprises the steps of:

comparing the level of at least one transcript in a first sample of a tissue to a second sample, wherein the first sample is of a tissue suspected of being neoplastic and the second sample is of a normal human tissue, wherein said transcript is identified by a tag selected from the group consisting of those shown Table 5;

identifying the first sample as neoplastic when the level of the at least one transcript is found to be higher in the first sample than in the second sample.

According to another embodiment of the invention a method is provided to aid in the determination of a prognosis for a colon cancer patient. The method comprises the steps of:

comparing the level of at least one transcript in a first sample of a tissue to a second sample, wherein the first sample is of a neoplastic colonic tissue and the second sample is of a normal human colonic tissue, and wherein the transcript is identified by a tag selected from the group consisting of those shown in Table 3; determining a poorer prognosis if the level of the at least one transcript is found to be lower in the first sample than in the second sample.

According to another aspect of the invention a method to aid in determining a prognosis for a patient with colon cancer is provided. The method comprises the steps of:

comparing the level of at least one transcript in a first tissue sample to a second sample, wherein the first sample is of a colonic cancer tissue and the second sample is of a normal human colonic tissue, and wherein the transcript is identified by a tag selected from the group consisting of those shown in Table 2;

determining a poorer prognosis if the level of the at least one transcript is found to be higher in the first sample than in the second sample.

In yet another embodiment of the invention a method is provided for diagnosing colon cancer in a sample suspected of being neoplastic. The method comprises the steps of:

comparing the level of expression of at least one protein in a first sample of a tissue to a second sample, wherein the first sample is of a colonic tissue suspected of being neoplastic and the second sample is of a normal human colonic tissue, and wherein the protein is encoded by a transcript identified by a tag selected from the group consisting of those shown in Table 3;

identifying the first sample as neoplastic when the level of expression of the protein is found to be lower in the first sample than in the second sample.

In another aspect of the invention a method of diagnosing colon cancer in a sample suspected of being neoplastic is provided. The method comprises the steps of:

comparing the level of expression of at least one protein in a first sample of a tissue to a second sample, wherein the first sample is of a colonic tissue suspected of being neoplastic and the second sample is of a normal human colonic tissue, and wherein the protein is encoded by a transcript

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identified by a tag selected from the group consisting of those shown in Table 2;

identifying the first sample as neoplastic when expression of the protein is found to be higher in the first sample than in the second sample.

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According to another embodiment of the invention a method is provided to aid in determining a prognosis of a patient having pancreatic cancer. The method comprises the steps of:

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comparing the level of at least one transcript in a first sample of a tissue to a second sample, wherein the first sample is of a neoplastic pancreatic tissue and the second sample is of a normal human colon tissue, wherein said transcript is identified by a tag selected from the group consisting of those shown Table 4;

of those shown Table 5;

determining a poorer prognosis if transcription is found to be higher in the first sample than in the second sample.

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In yet another aspect of the invention a method to aid in providing a prognosis for a cancer patient is provided. The method comprises the steps of:

comparing the level of at least one transcript in a first sample of a tissue to a second sample, wherein the first sample is of a neoplastic tissue and the second sample is of a normal human tissue of the same tissue type,

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determining a poorer prognosis if transcription is found to be higher in the first sample than in the second sample.

wherein said transcript is identified by a tag selected from the group consisting

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According to still another aspect of the invention, a method is provided for diagnosing pancreatic cancer in a sample suspected of being neoplastic. The method comprises the steps of:

comparing the level of expression of at least one protein encoded by a transcript in a first sample of a tissue to a second sample, wherein the first sample is of a pancreatic tissue suspected of being neoplastic and the second sample is of a normal human colon tissue, wherein said protein is

encoded by a transcript identified by a tag selected from the group consisting of those shown Table 4;

identifying the first sample as neoplastic when expression of the protein is found to be higher in the first sample than in the second sample.

According to yet another aspect of the invention a method is provided for diagnosing cancer in a sample suspected of being neoplastic. The method comprises the steps of:

comparing the level of expression of at least one protein in a first sample of a tissue to a second sample, wherein the first sample is of a tissue suspected of being neoplastic and the second sample is of a normal human tissue, wherein said protein is encoded by a transcript identified by a tag selected from the group consisting of those shown Table 5;

identifying the first sample as neoplastic when expression of the protein is found to be higher in the first sample than in the second sample.

In still another embodiment of the invention a method is provided to aid in the determination of a prognosis of a colon cancer patient. The method comprises the steps of:

comparing the level of expression of at least one protein in a first sample of a tissue to a second sample, wherein the first sample is of a neoplastic colonic tissue and the second sample is of a normal human colonic tissue, and wherein the protein is encoded by a transcript identified by a tag selected from the group consisting of those shown in Table 3;

determining a poorer prognosis if the level of expression is found to be lower in the first sample than in the second sample.

In still another embodiment of the invention a method is provided to aid in determining a prognosis for a patient with colon cancer. The method comprises the steps of:

comparing the level of expression of at least one protein in a first tissue sample to a second sample, wherein the first sample is of a colonic cancer tissue and the second sample is of a normal human colonic tissue, and

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wherein the protein is encoded by a transcript identified by a tag selected from the group consisting of those shown in Table 2;

determining a poorer prognosis if the level of expression is found to be higher in the first sample than in the second sample.

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In still another aspect of the invention a method is provided to aid in determining a prognosis of a patient having pancreatic cancer. The method comprises the steps of:

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comparing the level of expression of at least one protein in a first sample of a tissue to a second sample, wherein the first sample is of a neoplastic pancreatic tissue and the second sample is of a normal human colon tissue, wherein said protein is encoded by a transcript identified by a tag selected from the group consisting of those shown Table 4;

determining a poorer prognosis if the level of expression is found to be higher in the first sample than in the second sample.

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According to even a further aspect of the invention a method is provided to aid in providing a prognosis for a cancer patient. The method comprises the steps of:

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comparing the level of expression of at least one protein in a first sample of a tissue to a second sample, wherein the first sample is of a neoplastic tissue and the second sample is of a normal human tissue of the same tissue type, wherein said protein is encoded by a transcript identified by a tag selected from the group consisting of those shown Table 5;

determining a poorer prognosis if the level of expression is found to be higher in the first sample than in the second sample.

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In still another embodiment of the invention a method of treating a cancer cell is provided. The method comprises the step of:

administering to a cancer cell an antibody which specifically binds to a protein encoded by a transcript identified by a tag selected from the group consisting of those shown in Tables 2, 4, and 5, wherein the antibody is linked to a cytotoxic agent.

In another aspect of the invention an antibody linked to a cytotoxic agent is provided. The antibody specifically binds to a protein encoded by a transcript identified by a tag selected from the group consisting of those shown in Tables 2, 4, and 5.

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According to another aspect of the invention, a method of detecting colon cancer in a patient is provided. The method comprises the steps of:

comparing the level of at least one protein or transcript in a first

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body sample to a second body sample, wherein the first sample is a body sample of the patient and the second sample is of a normal human, wherein the protein is encoded by a transcript and the transcript is identified by a tag selected from the group consisting of those shown in Table 2, wherein the first and second body sample is a sample selected from the group consisting of blood, urine, feces, sputum, and serum;

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identifying neoplasia when the level of the at least one protein or transcript is found to be higher in the first sample than in the second sample.

In another aspect of the invention a method of detecting pancreatic cancer in a patient is provided. The method comprises the steps of:

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comparing the level of at least one protein or transcript encoded by a transcript in a first sample of a tissue to a second sample, wherein the first sample is of the patient and the second sample is of a normal human, wherein said protein is encoded by a transcript and the transcript is identified by a tag selected from the group consisting of those shown Table 4, wherein the first and second sample is a sample selected from the group consisting of blood, urine, feces, sputum, and serum;

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identifying neoplasia when the level of the at least one protein or transcript is found to be higher in the first sample than in the second sample.

Also provided by the present invention is a method of detecting cancer in a patient. The method comprises the steps of:

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comparing the level of at least one protein or transcript in a first sample to a second sample, wherein the first sample is of patient and the second sample is of a normal human, wherein said protein is encoded by a

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transcript and the transcript is identified by a tag selected from the group consisting of those shown Table 5, wherein the first and second body sample is a sample selected from the group consisting of blood, urine, feces, sputum, and serum;

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identifying neoplasia when the level of the at least one protein or transcript is found to be higher in the first sample than in the second sample.

Additionally provided by the present invention is a method to aid in the determination of a prognosis for a colon cancer patient. The method comprises the steps of:

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comparing the level of at least one protein or transcript in a first sample to a second sample, wherein the first sample is of a colon cancer patient and the second sample is of a normal human, wherein the protein is encoded by a transcript and the transcript is identified by a tag selected from the group consisting of those shown in Table 3, wherein the first and second body sample is a sample selected from the group consisting of blood, urine, feces, sputum, and serum;

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determining a poorer prognosis if the level of the at least one protein or transcript is found to be lower in the first sample than in the second sample.

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Provided by another embodiment of the invention is a method to aid in determining a prognosis for a patient with colon cancer. The method comprises the steps of:

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comparing the level of at least one protein or transcript in a first sample to a second sample, wherein the first sample is of a colonic cancer patient and the second sample is of a normal human, wherein the protein is encoded by a transcript and the transcript is identified by a tag selected from the group consisting of those shown in Table 2, wherein the first and second sample is a sample selected from the group consisting of blood, urine, feces, sputum, and serum;

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determining a poorer prognosis if the level of the at least one protein or transcript is found to be higher in the first sample than in the second sample.

According to still another aspect of the invention, a method to aid in determining a prognosis of a patient having pancreatic cancer is provided. The method comprises the steps of:

comparing the level of at least one protein or transcript in a first sample to a second sample, wherein the first sample is of a pancreatic cancer patient and the second sample is of a normal human, wherein said protein is encoded by a transcript and the transcript is identified by a tag selected from the group consisting of those shown Table 4, wherein said first and second sample is a sample selected from the group consisting of blood, urine, feces, sputum, and serum;

determining a poorer prognosis if the level of the at least one protein or transcript is found to be higher in the first sample than in the second sample.

Also provided by the present invention is a method to aid in providing a prognosis for a cancer patient. The method comprises the steps of:

comparing the level of expression of at least one protein or transcript in a first sample to a second sample, wherein the first sample is of a cancer patient and the second sample is of a normal human, wherein said protein is encoded by a transcript and the transcript is identified by a tag selected from the group consisting of those shown Table 5, wherein the first and second sample is a sample selected from the group consisting of blood, urine, feces, sputum, and serum;

determining a poorer prognosis if the level of the at least one protein or transcript is found to be higher in the first sample than in the second sample.

The present invention further includes antisense oligonucleotides complementary in whole or in part to SEQ ID NOS:1-732.

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This invention also provides a method for screening for candidate agents that modulate the expression of a polynuleotide selected from the group consisting of the polynucleotides in SEQ ID NOS.1-732 or their respective complements, by contacting a test agent with a pancreatic or colon cell and monitoring expression of the polynucleotide, wherein the test agent which modifies the expression of the polynucleotide is a candidate agent.

The present invention provides the art with new methods and reagents for diagnosing and prognosing cancers. In addition, some of the newly disclosed genes may play an important role in the development of cancers.

BRIEF DESCRIPTION OF THE DRAWINGS

Fig. 1. Comparison of expression patterns in colorectal cancers and normal colon epithelium. (FIG. 1A) A semi-logarithmic plot reveals 51 tags that were decreased more than 10 fold in primary CR cancer cells whereas 32 tags were increased more than 10 fold. 62,168 and 60,878 tags derived from normal colon epithelium and primary CR cancers, respectively, were used for this analysis. The relative expression of each transcript was determined by dividing the number of tags observed in tumor and normal tissue as indicated. To avoid division by 0, a tag value of 1 was used for any tag that was not detectable in one of the samples. These ratios were then rounded to the nearest integer and their distribution plotted on the abscissa. The number of genes displaying each ratio was plotted on the ordinate. Tu: CR tumors; NC: Normal colon. (FIG. 1B and FIG. 1C) Differentially expressed genes in The number of transcripts found to be differentially colorectal cancers. expressed (P < 0.01) are presented as Venn diagrams. Diagrams of transcripts that were decreased (FIG. 1B) or increased (FIG. 1C) in CR cancers compared to normal colon epithelium. Comparisons were between primary tumors and cells in culture as indicated.

Fig. 2. Northern blot analysis of genes differentially expressed in gastrointestinal neoplasia. Northern blot analysis was performed on total RNA (5 µg isolated from primary CR carcinomas (T) and matching normal colon epithelium (N), or pancreatic carcinomas. The top panel in each case show an

example of the ethidium bromide stained gels prior to transfer. The number of SAGE tags observed in the original analysis is indicated to the right of each blot. (FIG. 2A) Examples of transcripts that were decreased or increased in CR cancers. (FIG.2B) Examples of transcripts increased in pancreatic cancers (10). (FIG.2C) Examples of transcripts elevated in cancer which were or were not cancer type specific. Probes used for Northern blot analysis were as follows (Human SAGE Tag unique identifier, gene name, (GenBank accession number)): (FIG. 2A) H204104, Guanylin (M95714); H259108, (see Table 2); H1000193, (see Table 2); H998030, (see Table 2). (FIG. 2B) H294155, RIG-E (U42376); H560056, TIMP-1 (S68252). (FIG. 2C) H802810, EST338411 (W52120); H85882, 1-8D (X57351); H618841, GA733-1 (X13425).

Tables 2-5. Transcripts Differentially Expressed in Human Cancer.

Tag sequence represents the NlaIII site plus the adjacent 11 bp SAGE tag. Tag number indicates a SAGE UID (unique identifier). NC, TU, CL, PT, PC, refers to the number of the indicated tag observed in RNA isolated from normal colorectal epithelium, primary colorectal cancers, colorectal cancer cell lines, primary pancreatic cancers, or pancreatic cancer cell lines, respectively. The Accession and Gene Name refer to representative GenBank entries that contain the tag sequence.

Table 2 Transcripts increased in colorectal cancer.

Table 3 Transcripts decreased in colorectal cancer.

Table 4 Transcripts increased in pancreatic cancer.

Table 5 Transcripts increased in pancreatic and colorectal cancer.

DETAILED DESCRIPTION

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The inventors have discovered sets of human genes which are either upregulated or downregulated in cancer cells, as compared to normal cells. Specifically, certain genes have been found to be upregulated or downregulated in colorectal and/or pancreatic cancer cells, when compared to normal colon cells. These sets of differentially regulated genes can be used as diagnostic markers, either individually or in sets of, for example, 2, 5, 10, 20, or 30.

Genes whose expression was detected to be increased in colorectal cancer are shown in Table 2. Genes whose expression was detected to be decreased in colorectal cancer are shown in Table 3. Genes whose expression was detected as increased in pancreatic cancer are shown in Table 4. Genes whose expression was detected as increased in both pancreatic cancer and colorectal cancer are shown in Table 5. These latter genes likely play a role in neoplastic development generally.

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Tag sequences, as provided herein, uniquely identify genes. This is due to their length, and their specific location (3') in a gene from which they are drawn. The full length genes can be identified by matching the tag to a gene data base member, or by using the tag sequences as probes to physically isolate previously unidentified genes from cDNA libraries. The methods by which genes are isolated from libraries using DNA probes are well known in the art. See, for example, Veculescu et al., Science 270: 484 (1995), and Sambrook et al. (1989), MOLECULAR CLONING: A LABORATORY MANUAL, 2nd ed. (Cold Spring Harbor Press, Cold Spring Harbor, New York). Once a gene or transcript has been identified, either by matching to a data base entry, or by physically hybridizing to a cDNA molecule, the position of the hybridizing or matching region in the transcript can be determined. If the tag sequence is not in the 3' end, immediately adjacent to the restriction enzyme used to generate the SAGE tags, then a spurious match may have been made. Confirmation of the identity of a SAGE tag can be made by comparing transcription levels of the tag to that of the identified gene in certain cell types.

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In addition to the sequences shown in SEQ ID NOS: 1-732, or their complements, this invention also provides the anti-sense polynucleotide stand, e.g. antisense RNA to these sequences or their complements. One can obtain an antisense RNA using the sequences provided in SEQ ID NOS: 1-732 and the methodology described in Vander Krol et al. (1988) BioTechniques 6:958.

The invention also encompasses polynucleotides which differ from that of the polynucleotides described above, but which produce the same phenotypic effect, such as the allele. These altered, but phenotypically equivalent polynucleotides are referred to "equivalent nucleic acids." This invention also encompasses polynucleotides characterized by changes in non-coding regions that do not alter the phenotype of the polypeptide produced therefrom when compared to the polynucleotide herein. This invention further encompasses polynucleotides, which hybridize to the polynucleotides of the subject invention under conditions of moderate or high stringency.

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The polynucleotides can be conjugated to a detectable marker, e.g., an enzymatic label or a radioisotope for detection of nucleic acid and/or expression of the gene in a cell. A wide variety of appropriate detectable markers are known in the art, including fluorescent, radioactive, enzymatic or other ligands, such as avidin/biotin, which are capable of giving a detectable signal. In preferred embodiments, one will likely desire to employ a fluorescent label or an enzyme tag, such as urease, alkaline phosphatase or peroxidase, instead of radioactive or other environmental undesirable reagents. In the case of enzyme tags, colorimetric indicator substrates are known which can be employed to provide a means visible to the human eye or spectrophotometrically, to identify specific hybridization with complementary nucleic acid-containing samples. Briefly, this invention further provides a method for detecting a single-stranded polynucleotide identified by SEQ ID NOS.1-732 or its complement, by contacting target single-stranded polynucleotides with a labeled, single-stranded polynucleotide (a probe) which is at least 10 nucleotides of the complement of SEQ ID NOS: 1-732 (or the corresponding complement) under conditions permitting hybridization (preferably moderately stringent hybridization conditions) of complementary single-stranded polynucleotides, or more preferably, under highly stringent hybridization conditions. Hybridized polynucleotide pairs are separated from un-hybridized, single-stranded polynucleotides. The hybridized polynucleotide

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pairs are detected using methods well known to those of skill in the art and set forth, for example, in Sambrook et al. (1989) supra.

The polynucleotides of this invention can be isolated using the technique described in the experimental section or replicated using PCR. The PCR technology is the subject matter of United States Patent Nos.4,683,195, 4,800,159, 4,754,065, and 4,683,202 and described in PCR: The Polymerase Chain Reaction (Mullis et al. eds, Birkhauser Press, Boston (1994)) or MacPherson et al. (1991) and (1994), supra, and references cited therein. Alternatively, one of skill in the art can use the sequences provided herein and a commercial DNA synthesizer to replicate the DNA. Accordingly, this invention also provides a process for obtaining the polynucleotides of this invention by providing the linear sequence of the polynucleotide, nucleotides, appropriate primer molecules, chemicals such as enzymes and instructions for their replication and chemically replicating or linking the nucleotides in the proper orientation to obtain the polynucleotides. In a separate embodiment, these polynucleotides are further isolated. Still further, one of skill in the art can insert the polynucleotide into a suitable replication vector and insert the vector into a suitable host cell (procaryotic or eucaryotic) for replication and amplification. The DNA so amplified can be isolated from the cell by methods well known to those of skill in the art. A process for obtaining polynucleotides by this method is further provided herein as well as the polynucleotides so obtained.

RNA can be obtained by first inserting a DNA polynucleotide into a suitable host cell. The DNA can be inserted by any appropriate method, e.g., by the use of an appropriate gene delivery vector or by electroporation. When the cell replicates and the DNA is transcribed into RNA; the RNA can then be isolated using methods well known to those of skill in the art, for example, as set forth in Sambrook et al. (1989) supra. For instance, mRNA can be isolated using various lytic enzymes or chemical solutions according to the procedures set forth in Sambrook et al. (1989), supra or extracted by nucleic-acid-binding resins following the accompanying instructions provided by manufactures.

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Polynucleotides having at least 10 nucleotides and exhibiting sequence complementarity or homology to SEQ ID NOS: 1-732 find utility as hybridization probes. In some aspects, the full coding sequence of the transcript, i.e., for SEQ ID NOS: 1-732, are known. Accordingly, any portion of the known sequences available in GenBank, or homologous sequences, can be used in the methods of this invention.

It is known in the art that a "perfectly matched" probe is not needed for a specific hybridization. Minor changes in probe sequence achieved by substitution, deletion or insertion of a small number of bases do not affect the hybridization specificity. In general, as much as 20% base-pair mismatch (when optimally aligned) can be tolerated. Preferably, a probe useful for detecting the aforementioned mRNA is at least about 80% identical to the homologous region of comparable size contained in the previously identified sequences identified by SEQ ID NOS:1-732, which correspond to previously characterized genes or SEQ ID NOS:1-732, which correspond to known ESTs. More preferably, the probe is 85% identical to the corresponding gene sequence after alignment of the homologous region; even more preferably, it exhibits 90% identity.

These probes can be used in radioassays (e.g. Southern and Northern blot analysis) to detect, prognose, diagnose or monitor various pancreatic or colon cells or tissue containing these cells. The probes also can be attached to a solid support or an array such as a chip for use in high throughput screening assays for the detection of expression of the gene corresponding to one or more polynucleotide(s) of this invention. Accordingly, this invention also provides at least one of the transcripts identified as SEQ ID NOS:1-732, or its complement, attached to a solid support for use in high throughput screens.

The total size of fragment, as well as the size of the complementary stretches, will depend on the intended use or application of the particular nucleic acid segment. Smaller fragments will generally find use in hybridization embodiments, wherein the length of the complementary region may be varied,

such as between about 10 and about 100 nucleotides, or even full length according to the complementary sequences one wishes to detect.

Nucleotide probes having complementary sequences over stretches greater than 10 nucleotides in length are generally preferred, so as to increase stability and selectivity of the hybrid, and thereby improving the specificity of particular hybrid molecules obtained. More preferably, one can design polynucleotides having gene-complementary stretches of more than 50 nucleotides in length, or even longer where desired. Such fragments may be readily prepared by, for example, directly synthesizing the fragment by chemical means, by application of nucleic acid reproduction technology, such as the PCR technology with two priming oligonucleotides as described in U.S. Pat. No. 4,603,102 or by introducing selected sequences into recombinant vectors for recombinant production. A preferred probe is about 50-75 or more preferably, 50-100, nucleotides in length.

The polynucleotides of the present invention can serve as primers for the detection of genes or gene transcripts that are expressed in pancreatic or colon cells. In this context, amplification means any method employing a primer-dependent polymerase capable of replicating a target sequence with reasonable fidelity. Amplification may be carried out by natural or recombinant DNA-polymerases such as T7 DNA polymerase, Klenow fragment of E.coli DNA polymerase, and reverse transcriptase.

A preferred amplification method is PCR. However, PCR conditions used for each reaction are empirically determined. A number of parameters influence the success of a reaction. Among them are annealing temperature and time, extension time, Mg²⁺ ATP concentration, pH, and the relative concentration of primers, templates, and deoxyribonucleotides. After amplification, the resulting DNA fragments can be detected by agarose gel electrophoresis followed by visualization with ethidium bromide staining and ultraviolet illumination.

The invention further provides the isolated polynucleotide operatively linked to a promoter of RNA transcription, as well as other regulatory

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sequences for replication and/or transient or stable expression of the DNA or RNA. As used herein, the term "operatively linked" means positioned in such a manner that the promoter will direct transcription of RNA off the DNA molecule. Examples of such promoters are SP6, T4 and T7. In certain embodiments, cell-specific promoters are used for cell-specific expression of Vectors which contain a promoter or a the inserted polynucleotide. promoter/enhancer, with termination codons and selectable marker sequences, as well as a cloning site into which an inserted piece of DNA can be operatively linked to that promoter are well known in the art and commercially available. For general methodology and cloning strategies, see Gene Expression Technology (Goeddel ed., Academic Press, Inc. (1991)) and references cited therein and Vectors: Essential Data Series (Gacesa and Ramji, eds., John Wiley & Sons, N.Y. (1994)), which contains maps, functional properties, commercial suppliers and a reference to GenEMBL accession numbers for various suitable vectors. Preferable, these vectors are capable of transcribing RNA in vitro or in vivo.

Fragment of the sequences shown in SEQ ID NOS:1-732 or their respective complements also are encompassed by this invention, preferably at least 10 nucleotides and more preferably having at least 18 nucleotides. Larger polynucleotides, e.g., cDNA or genomic DNA, which hybridize under moderate or stringent conditions to the polynucleotide sequences shown in SEQ ID NOS:1-732, or their respective complements, also are encompassed by this invention.

In one embodiment, these fragments are polynucleotides that encode polypeptides or proteins having diagnostic and therapeutic utilities as described herein as well as probes to identify transcripts of the protein which may or may not be present. These nucleic acid fragments can by prepared, for example, by restriction enzyme digestion of the polynucleotide of SEQ ID NOS:1-732, or their complements, and then labeled with a detectable marker. Alternatively, random fragments can be generated using nick translation of the molecule. For

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methodology for the preparation and labeling of such fragments, see Sambrook et al., (1989) supra.

Expression vectors containing these nucleic acids are useful to obtain host vector systems to produce proteins and polypeptides. It is implied that these expression vectors must be replicable in the host organisms either as episomes or as an integral part of the chromosomal DNA. Suitable expression vectors include viral vectors, including adenoviruses, adeno-associated viruses, retroviruses, cosmids, etc. Adenoviral vectors are particularly useful for introducing genes into tissues in vivo because of their high levels of expression and efficient transformation of cells both in vitro and in vivo. When a nucleic acid is inserted into a suitable host cell, e.g., a procaryotic or a eucaryotic cell and the host cell replicates, the protein can be recombinantly produced. Suitable host cells will depend on the vector and can include mammalian cells, animal cells, human cells, simian cells, insect cells, yeast cells, and bacterial cells constructed using well known methods. See Sambrook et al. (1989) supra. In addition to the use of viral vector for insertion of exogenous nucleic acid into cells, the nucleic acid can be inserted into the host cell by methods well known in the art such as transformation for bacterial cells; transfection using calcium phosphate precipitation for mammalian cells; or DEAE-dextran; electroporation; or microinjection. See Sambrook et al. (1989) supra for this methodology. Thus, this invention also provides a host cell, e.g. a mammalian cell, an animal cell (rat or mouse), a human cell, or a procaryotic cell such as a bacterial cell, containing a polynucleotide encoding a protein or polypeptide or antibody.

When the vectors are used for gene therapy in vivo or ex vivo, a pharmaceutically acceptable vector is preferred, such as a replication-incompetent retroviral or adenoviral vector. Pharmaceutically acceptable vectors containing the nucleic acids of this invention can be further modified for transient or stable expression of the inserted polynucleotide. As used herein, the term "pharmaceutically acceptable vector" includes, but is not limited to, a vector or delivery vehicle having the ability to selectively target

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and introduce the nucleic acid into dividing cells. An example of such a vector is a "replication-incompetent" vector defined by its inability to produce viral proteins, precluding spread of the vector in the infected host cell. An example of a replication-incompetent retroviral vector is LNL6 (Miller, A.D. et al. (1989) BioTechniques 7:980-990). The methodology of using replication-incompetent retroviruses for retroviral-mediated gene transfer of gene markers is well established (Correll et al. (1989) PNAS USA 86:8912; Bordignon (1989) PNAS USA 86:8912-52; Culver, K. (1991) PNAS USA 88:3155; and Rill, D.R. (1991) Blood 79(10):2694-700. Clinical investigations have shown that there are few or no adverse effects associated with the viral vectors, see Anderson (1992) Science 256:808-13.

Compositions containing the polynucleotides of this invention, in isolated form or contained within a vector or host cell are further provided herein. When these compositions are to be used pharmaceutically, they are combined with a pharmaceutically acceptable carrier.

This invention further encompasses genes, either genomic or cDNA, which code for a polypeptide or protein in the cell of interest. The genes specifically hybridize under moderate or stringent conditions to a polynucleotide identified by SEQ ID NOS: 1-732 or their respective complements. The process of identification of larger fragment or the full-length coding sequence to which the partial sequence depicted in SEQ ID NOS:1-732 hybridizes preferably involves the use of the methods and reagents provided in this invention, either singularly or in combination.

Five methods are disclosed herein which allows one of skill in the art to isolate the gene or cDNA corresponding to the transcripts of the invention.

RACE-PCR Technique

One method to isolate the gene or cDNA which code for a polypeptide or protein and which corresponds to a transcript of this invention, involves the 5'-RACE-PCR technique. In this technique, the poly-A mRNA that contains the coding sequence of particular interest is first identified by hybridization to

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a sequence disclosed herein and then reverse transcribed with a 3'-primer comprising the sequence disclosed herein. The newly synthesized cDNA strand is then tagged with an anchor primer of a known sequence, which preferably contains a convenient cloning restriction site attached at the 5'end. The tagged cDNA is then amplified with the 3'-primer (or a nested primer sharing sequence homology to the internal sequences of the coding region) and the 5'-anchor primer. The amplification may be conducted under conditions of various levels of stringency to optimize the amplification specificity. 5'-RACE-PCR can be readily performed using commercial kits (available from, e.g., BRL Life Technologies Inc, Clotech) according to the manufacturer's instructions.

Identification of known genes or ESTs

In addition, databases exist that reduce the complexity of ESTs by assembling contiguous EST sequences into tentative genes. For example, TIGR has assembled human ESTs into a datable called THC for tentative human consensus sequences. The THC database allows for a more definitive assignment compared to ESTs alone. Software programs exist (give examples) that allow for assembling ESTs into contiguous sequences from any organism.

Isolation of cDNAs from a library by probing with the SAGE transcript or tag

Alternatively, mRNA from a sample preparation was used to construct cDNA library in the ZAP Express vector following the procedure described in Velculescu et al. (1997) Science 270:484. The ZAP Express cDNA synthesis kit (Stratagene) was used accordingly to the manufacturer's protocol. Plates containing 250 to 2000 plaques are hybridized as described in Rupert et al. (1988) Mol. Cell. Bio. 8:3104 to oligonucleotide probes with the same conditions previously described for standard probes exxcept that the hybridization temperature is reduced to room temperature. Washes are performed in 6X standard-saline-citrate 0.1% SDS for 30 minutes at room temperature. The probes are labeled with 32P-ATP through use of T4 polynucletoide kinase.

Table 2 - Transcripts increased in colon cancer

Transcripts increased in only colon primary tumors compared to normal colon (61 genes)

NC: Normal Colon TU: Colon Primary Tumor

CL: Colon Cancer Cell Line
PT: Pancreatic Primary Tumor
PC: Pancreatic Cancer Cell Line

					1	1	5	Aggreen	Gene Name
[Tog Seguence	Tag Number	Š	10	CL	2	긴]	
=	1 ag Seducine	H284749	612	755	411	191	333	F15516	H.sapiens mitochondrial ES1 sequence (1-1-12) nom
-	CATGCACCIAAIIGG	7022201	Ç	505	23.5	8	314	U35430	Human cytochrome c oxidase subunit III (COIII) pse
2	CATGTGATTTCACTI	H933/04	;		S	443	161	Z70701	H.sapiens mRNA (fetal brain cDNA c2 11).
~	CATGCCTGTAATCCC	H388150	455				1		H.sapiens HNF1-C mRNA.
					T	1		X71346	H.sapiens HNF1-B mRNA.
		11001001	203	537	82	4	æ	U09500	Human mitochondrion cytochrome b gene, partial cds
7	CATGCACTACTCACC	H291202	365	5	389	453	194	X66785	H.sapiens mRNA for transacylase (DBT).
~	CATGGTGAAACCCCA(U)	00/00/1						X17648	Human mRNA for granulocyte-macrophage colony-stimu
					Γ			U09087	Human thymopoietin beta mRNA, complete cds.
								009088	Human thymopoietin gamma mRNA, complete cds.
								U20770	Human metastasis suppressor (KAII) mRNA, complete
			1	Ę	,	g	F	W15552	2b91h11.s1 Soares parathyroid tumor Nb11PA Homo sap
٥	CATGGGCTTTAGGGA	H687915	7	3//2	٩	;		10000	2c05d03.s1 Soares parathyroid tumor Nb11PA Homo sap
L								22000	111407 -1 Home saniens cDNA clone 138925 5'.
								107900	y in the contract of the contraction of the contrac
	* * * OOTHER : * * * *	H130369	32	272	32	23	20	X89839	H.sapiens mitochondrial DNA for loop attachment se
~	CATGACITICCAAA	7000111	1	177	9	30	2	T11555	A 1486F Homo sapiens cDNA clone A 1486 sunitar to MI
∞		HY02424	3/2	210	,	20	01	T15773	IB1870 Homo sapiens cDNA 3'end similar to Human mi
6	CATGAGGGTGTTTTC	7/8C/1H	3 8	217	- 12	2 2	28	X12544	Human mRNA for HLA class II DR-beta (HLA-DR B).
2	10 CATGAGGTCAGGAGA(1)	H1//313						\$73483	phosphorylase kinase catalytic subunit PHKG2 homol
		2000111	124	104	[2	Ξ	2	X74301	H.sapiens mRNA for MHC class II transactivator.
	CATGTTGGCCAGGC1	H1023344						U28687	Human zinc finger containing protein ZNF157 (ZNF15
			1					U29119	Human leiomyoma LM-196.4 ectopic sequence from HMG
			1					U56236	Human Fc alpha receptor b mRNA, complete cds.
		31371011	6	18	=	4	64	W03751	za62h11.rl Soares fetal liver spleen INFLS Homo sa
2	12 CATGATCACGCCCIC	0104171	1	1				W03770	za63f10.r1 Soares fetal liver spleen INFLS Homo sa
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						-	Т	za42109.rl Soares Ietal IIver spiecii (141 Lo 110110 Su
13 CATGGGGGTCAGGGG	H696691	37	170	=	9	<u>م</u>	T12078	A730R Homo sapiens CDNA clone A730 similar to mice
				1	1	1	1,0,50	11 6401 brain cDNA 3"-rud GFN-007C04.
14 CATGGCTAGGTITAT	H641789	38	44	=	2	=	71010	Himan fetal brain cDNA 3'-end GEN-117E01.
				1	1,	1:	20000	Toknowo
15 CATGCCCCGTACATC	H350996	28	2	2	1	<u>۔</u> اع	100130	Uman fatal brain cDNA 3'-end GEN-007D07.
16 CATGAGTAGGTGGCC	H183018	<u>~</u>	=	7	=	1	170107	Times for kein cond 3'-end GEN-009C05.
					1	+	DS1052	Human Icial Dian China 2 and GEN-089501
							D52836	Human retai orain Colvo 3 -cito Octo- vovico:
COUTO A TOTOGOT . S	H388278	82	124	19	71	23	D83195	Human DNA for Deoxyribonucicase i precurson.
1) CAIGCLIGIAGICCE	H136465	64	121	28	24	15	D54113	Human fetal brain cDNA 3-end GEN-129803.
18 CATGAGACCCACAAC	H327364	49	101	35	7	40	F15796	H.sapiens mitochondrial EST sequence (102-23) from
19 CATGCATTIGIANTA	H874182	28	78	14	0	2		Meel frament of
ZU CAIGICCCCGIACCI	HK06582	23	73	8	9	19	Z59183	H.sapiens CpC Island DINA Belloille Machinery
21 CATGGCCAACCICCI	200011						D52905	Human fetal brain cDNA 5'-end GEN-091D11.
Little Control	1400574	Š	73	-	14	91	F16449	H.sapiens mitochondrial EST sequence (129-09) from
22 CATGGCCATCCCLII	P20400H	,,,	12	2	35	14	U06452	Human melanoma antigen recognized by 1-cells (MAK)
23 CATGTTGGTCAGGCT	H102/3/0	3	8	2 2	2	92		
24 CATGTCCTATTAAG	H881003	3 6	;		-		D51004	Human fetal brain cDNA 3'-end GEN-006D02.
25 CATGITACTIATACT	H991026	7	7	1	1	+	1 49057	Homo sapiens retinal fovea EST HFD010904 sequence.
		_		T			D51071	Human fetal brain cDNA 3'-end GEN-010E01.
				1	T	,		
26 CATGATGGCAGGAGT	H238755	=	45	- -	4	,,		
22 CATGCTAAGGGAGG	H461411	S	44	7	7	7		
2) CATGGGTGAGACACT	H713234	7	44	2	=	2	103592	Human ADF/A I'r dailsiocaec illicites, e che, e che fam
20 CATCACCTGTATCCC	H97078	9	42	17	2	32	X57352	Human 1-90 gene Holli Interestion 150562 5' simil
30 CATGCCAGTCCGCCT	H339302	0	39	0	-	0	H015/1	yjosedoni nomo septems con a clone 151846 S' simil
200000000000000000000000000000000000000							H03072	yjdogiz.ri Homo sapiciis czywa cione 34011
J. S.	H802810	-	37	0	-	0	T25155	EST/30 Homo sapiens Cond Cross 2-2011
SI CATOTAANTITIOSS	H993264	٥	37	2	3	5	DS0972	Human fetal brain cDNA 3 -end GEN-004AV3.
32 CATGLIAGCTIOLLI	2227711	-					D51211	Human fetal brain cDNA 3'-end GEN-U1/E08.
		-					D\$2162	Human fetal brain cDNA 3'-end GEN-069r'04.
		-					T23865	seq2012 Homo sapiens cDNA clone Cot13/4Ft-4HB3MA-3
OTTO COLOR	A5250AU	-	12	_	0	0	M32053	Human H19 RNA gene, complete cds.
33 CATGGCCACCCCIO	DAC 10011	· =	35	2	33	51	X67247	H.sapiens mS8 gene for ribosomal protein 58.
34 CATGTAATAAGUIG	11,00,00	: : -	2	~	-	14	T11939	A953F Homo sapiens cDNA clone A953 similar to Mito
35 CATGTACTGCTCGGA	17011011	:						

	0760361	٥	7	22	õ	4	T95857	ye42f01.s1 Homo sapiens cDNA clone 120409 3' simil
16 CATGGTGAAACCCA	H/33/49	`	1	:	+	1	Γ	za35b09.rl Soares fetal liver spleen INFLS Homo sa
			1	T				za63g03.rl Soares fetal liver spleen INFLS Homo sa
A A CA	H\$26210	9	56	=	~	3	X54195	Human line-1 element DNA, host sequence flanking i
CATGGAAACTGAACA							U29607	Human methionine aminopeptidase mKNA, complete cus
							H95100	yw57b10.rl Homo sapiens cDNA clone 236313 3 simil
A TO A CHITTING A A A	H131009	-	22	4	-	٥		220
OTOTO O	H\$55450	0	21	7	6	12	D29062	Human keratinocyte culva, cione vol.
CALGGACIGCGIGCG							D29563	Human keratinocyte cDNA, clone 713.
TOATOUTOAOTOTO	H863923	4	12	7	2	-	T03196	FB3B5 Homo sapiens cDNA clone r B3B3 3 end.
100100	H7916	2	20	2	2	_	Z57093	H.sapiens CpG DNA, clone 104a10, reverse read cpg.
CATGAAACIGIGGII							Z60184	H.sapiens CpG island DNA genomic Msel fragment, ci
							Z63649	H.sapiens CpG island DNA genomic Msel Iragment, cl
							W31349	2b95d06.s1 Soares parathyroid tumor NOHPA Homo sap
100000	H699051	0	61	0	0	0		400 000 11 4 011 11 V
CATGGGGGGGGG		2	16	-	٥	0	W31448	zb96h01.s1 Soares parathyroid tumor North A north Sap
CATGGTGCCCUIGCC							W47282	zc40b06.rl Soares senescent fibroblasts Nb113r 110mo
A HO	1.1600144	-	61	5	2	~	X71428	H.sapiens fus mRNA.
CATGGGGGGTAACTA							S62140	TLS=translocated in liposarcoma [human, mkNA, 1624
							W31782	zb96a06.r1 Soares parathyroid tumor North'A riottic sap
TAUUU	H883079	-	6	14	27	91	M24398	Human parathymosin niRNA, complete cds.
CATGICCIOCCCAI	13050011	c	2	0	0	0		
CATGAAGTGGCAAGA	11700759	٥	2	0	0	0	U33317	Human defensin 6 (HD-6) gene, complete cds.
CATGGGTATTAACCA	0.00071	<u>`</u>					M98331	Homo sapiens defensin 6 mRNA, complete cds.
T. 000 1.0	11504317	,	2	0	2	-	D32027	Human mRNA for T cell receptor V beta 14 CDR3, par
CATGGGCIACACCII		1	2	0	7	-	T11701	A 1225F Homo sapiens cDNA clone A 1225 similar to MI
CLLLC	H175870	-	2	0	0	0	D51783	Human fetal brain cDNA 5'-end GEN-051 G02.
CATUAGOOIGITICO	TAACTCH	0	=	0	2	. 0	D13138	Human mRNA for dipeptidase.
SO CATGCAAGGACCAGC	101777	1						Homo sapiens (clones MDP4, MDP7) microsomal dipept
		1						RDP=renal dipeptidase [human, kidney, Genomic, 357
O V O F V V	H950498	-	100	0	167	0	M10629	Human alpha-1 collagen gene, 3' end with polyA sit
CAIGIOGAAAIGACC	U210514	-	=	~	4	-	H11641	ym 17e04.s1 Homo sapiens cDNA clone 4/902.3 Simila
CATGATCCGCCIGCC	D152131	· -					R95667	yq51a09.s1 Homo sapiens cDNA clone 199288 3 simil
U V U V II	HR75782	-	=	0	0	-		-
S3 CATGICCCGIACAC	H241665	0	=	0	12	14	M74090	Human TB2 gene mRNA, 3' end.
S4 CAIGAIGIAAAAAI								

101801 Himan ivsozyme mRNA, complete cds with an Alu tepe	1	M19045 Human lysozyme mRNA, complete cds.	ı		Vegas Human 1-8D gene from interferon-inducible gene fam	1331 Human - C. Berrera	X02490 Human interferon-inducible mRNA (cDNA 1-8).	1			102040 Human SPARC/osteonectin mRNA, complete cds.	٦	Tiesaia Human RNA fragment from patients with Crohn's dise	I		Alamo Alda Carit	Models Human chaperonin-like protein (HTKS) intrict, complete	-1	1 27706 Human chaperonin protein (1 cp.20) gene complete cus	٦	
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				H337744	110011	H85882				H165175	2112	T241747	11.65711	H310975		H613862		H992010			
				し上づつつつつでは、い	SS ICA ICCCACCCCOTC	LUCTULTOCOCCE	S6 CATGACCATTCTOCT			JULY JULY JULY JE	S7 ICA CACCACCA ICCC	(V)LUVUT OECH	S8 ICATGATGTGAAGAGT(A)	TOTTOCTTOCT	SY ICAICCACHIOCHICE	A TOTATOTOTOTO OF	CALCOCCITCIOCCA	A CONTACT A CONTRACT A	61 CATOLIAUATANCO		

19 CATGCTCCTCACCTG

Transcripts increased in both colon primary tumors and colon cancer cell lines compared to normal colon (47 genes)

NC: Normal Colon

TU: Colon Primary Tumor CL: Colon Cancer Cell Line PT: Pancreatic Primary Tumor PC: Pancreatic Cancer Cell Line

				_		_	1	_		-		_	_	_		_	Т	Τ	Т	_	1	Т	7	Т	Т	Т	_	-		1
uman ribosomal protein L28 mRNA, complete cds.	II mDNA for I Ren3	Hullian mixed to both	H.sapiens BBC1 mining	H. sapiens mRNA for 23 kD highly basic protein	H sapiens mRNA for elongation factor 2.		H.sapiens S19 ribusonilal process in the sapiens S19 ribu	Human acidic ribosomal phospholocini i z missy;	H.sapiens hng mRNA for uracil DNA glycosylase.	Human glyceraldehyde 3-phosphate dehydrogenase mKN	In maint many for elongation factor-1-gamma.	The separate state of the separate of the sepa	Human pancreatic tumor-related process since the	H. sapiens mRNA for ribosomal protein L8.	111 DNA for ribosomal protein L3.	H.Sapiens mixture for the sale of the sale	Human novel gene mixira, withplace cast	Human Wilm's tumor-related protein (QM) mKNA, comp	Ilaminin recentor homolog (3' region) [human, mRNA	DAIA 62 OPE	H.Sapiens minita for Old	Human mRNA for noosomal protein L32	Human ribosomal protein S4 (RPS4X) isoform mKNA, C	Human scar protein mRNA, complete cds.	H.sapiens mRNA for ribosomal protein S18.	Homo sapiens 18S ribosomal protein (HKE3) mRNA seq	Liman mana for T-cell cyclophilin.	Tr. DNA for insulin-like growth factor II (IGF-2);	nument Division in anniete of	Human Bak mining, complete cas.
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1	87	52	╀	+	4	42	ő	╀	+	\$		51	+	1	9	36	1	+	1		48	×	;	7		42		28	0	2
Number	H599350	H239533		H333689	H171113	H148949	ACCC0311	F7/70CU	H0/1034	H807748		1050408	27,77		HS5227	1090991		H174037			1744683	COOLL	H932680	H861056		H965603		H379369	\$18912	1782584
-	Т	CATOCCACCATAT	2 CATGATGGCTGGTA1	1 CATGCCCGTCCGGAA			S CATGAGCACCICCAG	6 CATGCTGGGTTAALA	7 CATGGGATTTGGCCT	Т	┰	7			A DUTTOTO A TOTO S.	10 CAIGAAICCIGIGGE	11 CATGGGACCACIGAR	12 CATGAGGGCTTCCAA				13 CATGAAGGTGGAGGA	14 CATGTGCACGTTTTC	15 CATGTCAGATCTTTG		IN CATGTGGTGTTGAGG		TABOLIOCTAC	1/ CAIOCCIAGOIGO	18 CATGCTIGGGIIIIG
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D14530 Human homolog of yeast ribosomal protein S28, comp		Human mRNA for ribosomal protein L37, complete				M24194 Human MHC protein homologous to chicken B complex	U14967 Human ribosomal protein L21 mRNA, complete cds.	X55954 Human mRNA for HL23 ribosomal protein homologue.	Γ		H71935 ys15f12.rl Homo sapiens cDNA clone 214693 3.	Z43914 H. sapiens partial cDNA sequence; clone c-10003.	T48545 hbc3221 Homo sapiens cDNA clone hbc3221 Send.	X04347 Human liver mRNA fragment UNA binding protein Ori	X00910 Human mRNA for IGF-11 precursor (insulin-line grow	X61156 H.sapiens mRNA for laminin-binding protein.	103799 Human colin carcinoma faminin-binding protein mKNA	U02032 Hunan ribosomal protein L23a mRNA, partial cds.	U14970 Human ribosomal protein S5 mRNA, complete cds.	Г	M36981 Human putative NDP kinase (nm23-H2S) mKNA, complet	L16785 [Homo sapiens c-myc transcription factor (put) inicinal				П	\neg	\neg	T	7	D28137 Human mRNA for BS1-2, complete cus.		\neg	X72718 H.saptens DNA for orphan ICA V-Deta segment (2005)
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T. A. C. T. C.	\exists			23 CATGACAICAICGAI	24 CATGTICAAIAAAA	25 CATGGAACACAICCA	26 CATGTTATGGGATCI	27 CATGGCATAATAGGI	28 CATGATTCTCCAGTA		29 CATGACTCCAAAAA			_	\neg	31 CATGIACAAAAICUA	32 CATGGAAAAATGGTT		33 CATGAAGAAGATAGA	34 CATGCCTTCGAGATC	35 CATGACTGGGTCTAL		_	36 CATGCAGCICACIUA	_		38 CA1001000000000000000000000000000000000	39 CATGGTTCACATTAG	40 CATGLGAAATAAAAC	41 CATGAAAAAAACII	42 CATG IGC I GCC I G I I		V C C C C C C C C C C C C C C C C C C C	43 CATOCTOATOCCAOA

147970 Clone 147970	Soares fetal heart North I w notice sapicals controlled	3.	Tect 176663 Colon carcinoma (Caco-2) cell line II Homo saplens		A A 3 O S S 8 O C D N A S end	04)(Vergit Himan mRNA for actin-binding protein (illamin) (AB		Vocati Himms mRNA for fibronectin (FN precursor).	I umini	1 226206 In cariene isoform I gene for L-type calcium channe	11.3aplons society	
		7 H121311			A A 205580	ההההעע	ALACAV	21100		10/704	206266	505077	
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		1121311	1171711					1 78 19 82 17 19 84 17 17 17 17 17 17 17 1	1101010	0 901000	1777100	0 0 10 10 0 0	
		# C # C # C # C # C # C # C # C # C # C							- 45 ICATGGCCCAAGGACC	Г	46 ICATGATCTTGTTACT	- COROCIETE	A TO A TO A A CIT COLOR

cell lines compared to normal colon (181 genes) Transcripts increased in only colon cancer

NC: Normal Colon
TU: Colon Primary Tumor
CL: Colon Cancer Cell Line
PT: Pancreatic Primary Tumor
PC: Pancreatic Cancer Cell Line

Cone Nome			S Human ribosomal protein S12.	Т	Т			Г	T	Т	Т	T		٦		14 Human ribosomal protein S16	Т	T		П			Г	Γ	Т	Т		Т	Т	47 Human territin L chain
	Accession	X16869	X53505	20001	C00717	L19739	X83412	Z32564	0819/X	0779011	27.000	17800	S64030	T91925	66999X	M60854	140230	0074M	X65/8	U14968	X79234	103537	U58682	V57810	1	4	4	C7/57W		M11147
	2	412	25		22	179	102				1		358	0	252	108	•	•	-	142	49	120	5	┿	+	-+-	=	2	_	145
	PT	136	+-	-	-	74	99	╀	1	4	1		104	5 48	2 55	┿			38	2 65	1 53	╄	╀	+	+	+		80	4	78 92
	7	+-	+-	-+	245	201	186	┿	1	1	4		8 179	176	172	┿	+	-	7 133	3 112	=	+-	╀	┿	+	-	41 89	30	_	49
	T	2	+-	-	7 83	53	╀	╌	+	4	\dashv	_	115 128	0	19 05	┿	+	80	30 37	38 53	+-	╁	+-	+	-+	39 4	27 4	2	_	34 4
	UN I	,†	\dagger	72	137	63	-	7	+	4	4		F	╁╴	f	1		6	-	T	t	\dagger	\dagger	+	7	-	-		_	
	Too Number	3C00001	C700/6H	H615043	H263478	H278636		Ē					H1027448	H906438	1133070	(1/2001)	H374027	H696375	H41531	11567488	V03VCV11	1604741	H01017	H549145	H857362	H416106	H475448	H955718		H359102
		Tag Sequence	CATGTGTTGAGAG	CATGGCGAGGAAGG	CATCOAACCATCCA	т		CATGAAAAAAAAA						- 1	CAIGICICCAIACC		Т	Τ		\neg			14 CATGGCCGTGTCCGC	15 CATGGACGACACGAG	1	\neg	7	\neg	_	TOTTO COTTO TOTAL
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retrocka	\vdash	H150997	0	0	77 (0	H	П	yl96f11.r1 Homo sapiens cDNA clone 45943 5.
CATGAGCATCTCCAG		_		-		_	7Z	Z44640 I	H. sapiens partial cDNA sequence; clone c-20003.
	-	1	\dagger	+	+		z		yz29e01.rl Homo sapiens cDNA clone 2844/12 5.
CATCCCTGTATGAG H621369 24	369	7	4	32	77 3	\vdash	\square	П	Human ribosomal protein 524 mKNA.
-	624	33	_	39	76 2	21 67	4	X53777	Human L23 mkNA to purative treeses PNA clone 650651 3' similar to
		27			74	23 87		AA223340 g	gb AA22334U AA2234U numbaphun separa
H677342	\dagger	l R	┿	ļ	72 2	27 61		U12404	Human Csa-19
+	+	<u>=</u>	┿	42	70	32 146			H.sapiens EST sequence (135-18) from skeletal muscle
H26	19	58	\vdash	46 (69	-			Homo sapiens macrophage migration initionaly recon
CATGCCAGACAGAC H335945 23		23		_	-	-+	\perp	7	H. Sapiens Tibosolinal protein Care.
	Н	7		\dashv	-		4	Т	Human transferorase (11x1)
CATGGTGTTAACCAG H769045 16		91	_		+	-+	4	Т	luman flooronia protein 1 38
-		6	-	13	-	}		Т	H.sapiens ribosomai protein 1530.
-	019	15	-	27	┪	-	4	Т	fuman class of glutannone of amore and a company of the company of
CATGGTTCCTGGCC H775658 31		31		76	63	-	4	Т	H.Sapiens lau mono.
	\vdash	32		88	, 79	42 6	× 89		H.sapiens KPS.20
_	Н	7		4	8	5	39	\neg	2c45e11.rl Soares seriescent librorians 103101
	\dashv			+	+	-+	4	Т	Limes have I mRNA for high mobility group protein I.
1	-	=			\dashv		4	Т	Human ribosomal protein S29
H200576		=		77	┽	╌	4	014910	Uman YPIPO ribosomal protein S3 (rpS3)
- H348756	┪	<u>~</u>		53	2			Т	Home caniens ribosomal protein L18 (RPL18)
H667269	\dashv	~		<u></u>	\$ s		4	L11300	J87301 rl Homo sapiens cDNA clone 44932 5.
A H786433	\dashv			∞ ;	\$ \$ \$	_	8 5	TU0230	H sanjens ribosomal protein S13.
H769605	\dagger	(۵		7 7	2 5	; - ; -	\bot	131657	Human unknown protein mRNA, partial cds.
CATGGCCAGCCAGC H608393 8	\dagger	<u>-</u>		 	+	┿	╀	H41030	yn92a10.r1 Homo sapiens cDNA clone 175866 5'.
HK85384 14	\dagger	4	_	24	47	23	15	M16660	Human 90-kDa heat-shock protein
H851981	+	P	Т	0	8	7	0	N57419	yw82e04.r1 Homo sapiens cDNA clone 238/30 3 simil
H583571	\dagger	9		22	46	27	81	X59357	Human mRNA for Epstein-Barr virus small KNAS (EBER)
\dagger	\dagger	1	Т		T	-	-	L21756	Homo sapiens acute myeloid leukemia associated protein
		\downarrow	7		1	\vdash		D17652	Human mRNA for HBp15/L22, complete cds.
\$10191	+	-	-	-	46	47	53	M64716	Human ribosomal protein S25
2761611	3113	-) «	, 1%	\$	╁	63	L06498	Homo sapiens ribosomal protein S20 (RPS20)
CATGGCTTTTANGOA HSSS33	H58533	+-	12	2	44	H	27	M61831	Human S-adenosylhomocysteine hydrolase (AHCY)
		┨							

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		2320311			37	0	\vdash	M33680	Human 26-kDa cell surface protein TAPA-1
77	CATGCTAAAAAAAA	H408/00	•	╬	╁	4	╀	Т	Homo sapiens dbpB-like protein
78	CATGGGGTTTTTATT	H704500	4	╬	+	╁	4	T	Human translational initiation factor 2 beta subunit
5	CATGCCGATCACCGG	H363799	1	+	+	╁	1	Т	29221 J. J. Soares fetal lung NbHL 19W Homo sapiens
8	CATGCCACAAGAAGA	H594051	٥	,	╀	+	1	Т	Human HL60 3'directed Mbol cDNA, HUMGS01477, clone
				\dagger	\dagger	+		Τ	Soares fetal lung NbHL19W Homo sapiens cDNA clone 303055 31.
				\dagger	+	+	1	Г	yv84c07.s1 Homo sapiens cDNA clone 249420 3' similar to contains Alu
							_ 	H83884	repetitive element;
	O V O O O TO TO TO TO TO	H908373	1	=	78	11 13	Ш	П	H.sapiens CDEI binding protein mRNA.
∞	CAIGICICIACCAC					_	ור		Homo sapiens amyloid protein homologue mrana, compi
				-	-	-	1		Human binding protein mKNA, partial cds.
							S	\Box	APPH=amyloid precursor protein homolog (numan, pla
5	CATOCITITION AG	H783697	-	0	25	3		П	zb06f02.r1 Soares fetal lung NoHL19 w Holing Sapieus
72	CATOOTTICCCCAAC						7	Т	yx36f06.rl Homo sapiens cDNA clone 203043 5
				-			_	\neg	yx62a03.rl Homo sapiens civity civile cocco
	JOD V JOLEGE CONTRACTOR OF THE PARTY OF THE	H388426	2	<u>س</u>	25	3 1	13 2		H. sapiens partial cDNA sequence; cione certacos:
8	CATOCCIOICCAGC			\vdash	-	-	>	W02723	zc65c03.s1 Soares fetal heart North 19 W molino sapiens
				T		-		N24893	yx99h09.s1 Homo sapiens cDNA clone 209921 3.
				1	+	\vdash		N32178	yy25b09.s1 Homo sapiens cDNA clone 2/2249 3.
		11055501	6	2	22	150	7	H21873	yl34b10.s1 Homo sapiens cDNA clone 160123 3' simil
8	CATGTCATCIGA	Недолог	1	:		-	F	H26394	yl48e12.s1 Homo sapiens cDNA clone 161518 3' simil
				T	-	-		H69857	yr88d02.s1 Homo sapiens cDNA clone 212355 5' simil
				1	\dagger	\vdash		H70714	yu69b11.s1 Homo sapiens cDNA clone 239037 3' simil
		11358781	1	_	22	16	31	X55110	Human mRNA for neurite outgrowth-promoting protein
85		07000	1	, -	77	╀┈	┞	X03168	Human mRNA for S-protein.
98	CATGGCCGGGCCCTC	H01/040	1	+	1	+	+		2032d09.s1 Stratagene colon (#937204) Homo sapiens cDNA clone 588595
		111023233	,		24.	7	2 A	AA143561	3' similar to contains LTR7.tl LTR7 repetitive element
87	CATGTTGCTCAAAAA	U102333	1	1	1	\dagger	╁		2001g11.s1 Stratagene colon (#937204) Homo sapiens cDNA clone 300400
							_	AA152342	3' similar to contains LTR7.t3 LTR7 repetitive element;
						╁	\vdash		2/86h11.51 Stratagene colon (#937204) Homo sapiens cDNA clone 311337
					_		_	AA115727	3' similar to contains LTR7.t1 LTR7 repetitive element
_	_	H262987	9	2	24	5	15	R76502	yi61f09.r1 Homo sapiens cDNA clone 143/33 3.
8	CALGCAAAICAGGA						-	T32681	EST52915 Homo sapiens cDNA 5' end similar to None.
			1			-		T34662	EST72468 Homo sapiens cDNA 5' end similar to None.
		SLPELSIT	上	~	23	4	7	H04634	yj49h03.rl Homo sapiens cDNA clone 132111/ 5.
8	CATGGAAGATGIGGG	ההרננום	4	.]		1			

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			t	+	-	-	-	F00364	H. sapiens partial cDNA sequence; clone 76D12; ver
			1	+	+		4	7	vio 1c05 s.1 Homo sapiens cDNA clone 149384 3'.
8	CATGGTGCTCATTCA	H761150	3	, 	3		+	Т	yv86c02.s1 Homo sapiens cDNA clone 249602 3' simil
1			1	+	+	-	L	Ī.	yv88f07.s1 Homo sapiens cDNA clone 249829 3' simil
7	OTITO CITITO COLO	H654464	4	5	23	2	2	138961	Homo sapiens putative transmembrane protein (B5)
	CATGCCTTTACTTG	H1046401	9	=	23	<u>-</u> 02	01	J04026	Human thioredoxin (TXN) mRNA
┰	CATOTITICIONAL	H1023250	-	4	22	0	4	D1 1078	Human RGH2 gene.
\neg	CATOLITICACACA	H\$89267	0	0	22	0	61	XS3279	Human mRNA for placental-like alkaline phosphatase
3 2	CATOGALITETERS	H166539	7	5	22	2 4	4		Human pyrroline 5-carboxylate reductase mRNA,
	CATOCCTTAACCTGG	H651359	2	4	22	2 4	4		Human glutamate dehydrogenase
2 2	CATOCTCTTCGAGAA	H490889	4	∞	22	27	19	П	Human mRNA for glutathione peroxidase
Т	CATGAGAACAAAACC	H132098	F	7	21	6	9		H.sapiens mRNA for proliferation-associated gene
_[CATOCCAGGAGAA	H346761	3	~	21	2 2	24		Human stimulator of TAK KNA binding (SKb)
-1						-		D16933	Human HepG2 3' region cDNA, clone nmd4111.
9	SUPPLIES A TOTA A A A A A A A A A A A A A A A A	H294155	0	_	20	47 10	107	U42376	Human retinoic acid induced RIG-E
3	TO CALCACACACACACACACACACACACACACACACACACA	H631331	7	~	20	4			Unknown
5	CATOCACACACAC	1-1989024	4	-	20	3 2	22	F17524	H.sapiens EST sequence (012-12-32) Irom skeicial in
2	CATOTOTOTO	H122449	4	-	2	<u>س</u>	7		Unknown
3	CATGACICIOCCAAG	H861095	<u> </u> -	5	9	. 21	1	W52942	zc03h05.r1 Soares parathyroid tumor NbHPA Homo sap
3		91662911	E	~	2	2	3	R21316	yg48h11.r1 Homo sapiens cDNA clone 35917 5' sumla
<u>6</u>		1951917	0	0	0	0	0	X00566	Human lipoprotein apoAl.
3		700yacri	6	~	6	5	5	M80244	Human E16 mRNA
5	_	170004	, c	ءا،	<u>~</u>	+	2	H27927	y158c11.s1 Homo sapiens cDNA clone 162452 3' simil
2		H00/310	, ,	, ~	 	╀╌	20	X57959	H.sapiens ribosomal protein L7.
2	CATGATTALLITICI	1420800	,	1	82	╁╌	15 A	AA299898	EST12509 Uterus tumor I Homo sapiens cDNA 5' end
2	CATGGACCCIOGGA	H686319	, [. ~	<u>∞</u>	╁	-	_	Human glycyl-tRNA synthetase
= :		H855049	1	2	<u>∞</u>	4	4	X76013	H.sapiens QRSHs mRNA for glutaminyl-tRNA synthetas
<u> </u>		H11785	0	7	=	0	~	W16529	zbi0ali.ri Soares fetal lung NbHL19W Homo sapiens
=	CALCANACIONACA					\vdash	-	W35192	zc70b05.r1 Soares fetal heart NbHH19W Homo sapiens
					T	\vdash	\vdash	W52451	zc45d09.rl Soares senescent fibroblasts NbHSF Homo
	A A OTTO COO A COST A COST	H288373	-	-	12	0	3	D38251	Human mRNA for RPB5 (XAP4)
-	CATOCACCCCCC	H28872	<u> </u> -	9	=	2	31	D52570	Human fetal brain cDNA 5'-end GEN-081G12.
2	IIS CALGAACIAAIACIA		1			\vdash		D52758	Human fetal brain cDNA 5'-end GEN-087A08.
			_		Γ		-	D55953	Human fetal brain cDNA 5'-end GEN-407H12.
	A COTO A TOTOCT .	H504187	-	0	=	2	9	M22490	Human bone morphogenetic protein-2B (BMP-2B)
9	116 CAIGCIGIACCIGGA	,,,,,,,							

	П	Π			Г		Γ	Τ	Π	Т	R80990 yi94c02.r1 Homo sapiens cDNA clone 146882 5'			П	S85655 Human prohibitin	M38188 Human unknown protein from clone pHGR74 mRNA, comp		D83174 Human collagen binding protein 2.	X70940 H.sapiens clongation factor I alpha-2.	T30623 EST 19638 Homo sapiens cDNA 5' end similar to None.	HUMGS0004747, Human Gene Signature, 3'-directed cDNA	C01011 sequence.	zm62d06.s1 Stratagene fibroblast (#937212) Homo sapiens cDNA clone	AA111865 530219 3'	W56516 zd16c08.rl Soares fetal heart NbHH19W Homo sapiens	H30299 yo77d04.rl Homo sapiens cDNA clone 183943 5' simil			W04495 za58b10.r1 Soares fetal liver spleen INFLS Homo sa	W23528 zc71g11.s1 Soares fetal heart NbHH19W Homo sapiens		X75598 H.sapiens nm23H1 gene.	T35470 EST85850 Homo sapiens cDNA 5' end similar to None.	T35536 EST86951 Homo sapiens cDNA 5' end similar to None.
M12529	X16539	M27691	M86667	X53743	226	Z26328	1122055	╀	Ļ	N42	R80	R95	F16	T50	_	┞	Y00711	_	_	_		<u>5</u>	_	AAI	WS	1 H3(HS	14 W0	W0	W2	12 DI	\vdash	0 T3	F
0	╀	╀	٣	0	╄	╀	╀	+	+	+	4	_	7	_	91	0	0	3 5	2	=	-		-		\vdash	2	\vdash	9	-	\vdash	9	5 11	┞	H
7 48	╀	╀	16	-	╁	╀	╀	╀		+	15 2		15 8		15 0	╀	15 0	15 23	╁	15 3	\vdash	_	\vdash		-	4	-	4	\vdash	\vdash	4	14	┝	H
17	+	+		╀		╁	+	+	╁	+	2	\vdash	9	-	- - -	╀	╀	2	╀	2	╀		\vdash		\vdash	-	\vdash	4		\vdash	9	╀	╀	H
2 6	+	-	٦	╀	╀	╫	+	- <u>-</u>	+	+	0	\vdash	-	╁	-	-	╀	-	0	╀╌	╁		\dagger		\dagger	 	\vdash	-		T	6	0	-	
1308663		H819213	T98867H		1786	1700077	/00077H	H/62554	1,170/11		H561787		H633002		H756407	H524541	H577840	C19551H	H910430	H18469	200					H980130		H822331			H\$08767	H673954	H075194	
	117 CATGCGACCCCACGC	118 CATGTAGAAAAAAA	OO V V V Children	119 CATGAICHIGAAAGG		_		_	123 CATGGTGGACCCCAA		_	124 CATGOAGCAGCTGGA	TOOON	ולט כאוסקרמקמעמים		126 CATGATIGGCTIAAA	127 CATGGAAAAATTAA	LA CATOOATCACAOTT	129 CATGAGCCIIIGIIG	- 1	131 CATGAACAGAAGCAA					JON DO V JET DE DE CET	וזל כאוחוסווראים	COLUMN ATOT COLUMN	ואו השואטאוטואטאומניי			134 CATGCTTAATCCTGA	135 CATGOOCAGAGGACC	136 CATGTOACTOAAGC

		t	}	+	-	725545	FET87066 Homo sapiens cDNA 5' end similar to None.
		1	-	+	-		vi33e11 e1 Homo sapiens cDNA clone 150596 3'.
CATGGATAGTTGTGG	H576495		+	<u>-</u>	- -	Т	2b17d08.s1 Homo sapiens cDNA clone 302319 3'.
			+	+	╀	Π	za92h06.s1 Homo sapiens cDNA clone 300059 3'.
	178887H	-	4	5	6 3	H90469	yv01e06.rl Homo sapiens cDNA clone 241474 5' simil
CATGGTGGTGGACAC			╀		\vdash	R76765	yi63g01.rl Homo sapiens cDNA clone 143952 5' simil
			+	+	┞	T35045	EST79335 Homo sapiens cDNA similar to None
THU THU COUNTY	H961304	0	9	<u>□</u>	2	HS1447	yo31a05.rl Homo sapiens cDNA clone 179504 5'.
CATGTGGGGIACCII	1301301	1	╀	╀	-	W46469	zc32c05.rl Soares senescent fibroblasts NbHSF Homo
			t		L	W51800	zc48e04.rl Soares senescent fibroblasts NbHSF Homo
			+	+	\vdash	П	yh77f08.r1 Homo sapiens cDNA clone 135783 5.
TAATA TATA	H1001313	F	9	2	<u> </u> 2	L	Human prothymosin-alpha
140 CAIGITCALIAIANI	H515821	. 0	↓_	╀	8 12	D80012	Human KIAA0190 protein
CAIGCIICIGIGIAC(I)	1125315	-	╀	-	2 5	U02389	Human hLON ATP-dependent protease mKNA
CATGACTGGCGAAGT	01007111			-	-	T29819	EST96617 Homo sapiens cDNA 5' end similar to A 1 P-d
	200750311	-	 	2	9	X14850	Human histone H2A.X.
CATGGAAAGAGCTGA	C6407CH	-	\	1 =	7	104088	Human DNA topoisomerase II (top2) mRNA
CATGCAACTCTATGG	C//697H	9 6	- -	: =	0	╀-	Human beta globin retrovirus-like repetitive element
CATGAAATTIGGIGG	COCOLL		,	+	-	1188396	EST28e05 Homo sapiens cDNA clone 28e05
1,0	11406114	1-	1	-	~	╀	H.sapiens p85Mcm mRNA.
CATGCTGCACTIACT	11000	1	+	+	\vdash	D28480	Human mRNA for hMCM2, complete cds.
			+	\dagger	-	D55716	Human B lymphoma mRNA for P1cdc47, complete cds.
	162120	6	1	=	-	T30327	EST14849 Homo sapiens cDNA 5' end similar to None.
CATGAATATTGAGAA	H35167	<u>.</u>	,	+	-	T34394	EST66942 Homo sapiens cDNA 5' end similar to None.
			+	t	\vdash	T47475	yb 14c03.r1 Homo sapiens cDNA clone 71140 5.
			+	T	\vdash	T50289	yb14h08.r1 Homo sapiens cDNA clone 71199 5'.
	H890535	0	-	13	2		Unknown
	H697495	0	7	=	2 7		Unknown
49 CA 100000000000000000000000000000000000	H329737	0	٥	12	4 4		Human inducible poly(A)-binding protein
150 CATGCCAAGAAAA	H1048113	0	~	12	4	12 D16891	Human HepG2 3' region cDNA, clone nmazci I.
1SI CATGITITIONIANA	H977034	0	0	12	0	0 M29882	Human apolipoprotein A-II
CATCICACOCATAG	H345789	0	~	12	7	4 249216	H. sapiens mitoxantrone-resistance associated mr.n.A.
155 CATGCCCACGGTTAG	H63325	0	-	12	-		Unknown
CATGAATTCCCCC	H548203	0	0	12	0	0	Unknown
155 CATGGACCICCGGGC	H921067	0	2	=	-	8 M93651	Human set gene
156 CATGIGAAICIOGOI	.,,,,,,,,,						

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H12992 0 1 10 6 3 D53402 H12992 0 1 10 6 3 D53402 T61971 H12913 0 0 10 1 2 T35761	TAGTG	H242380	0	~	2	6	7	DSS671	Human heterogeneous nuclear ribonucleoprotein
H12992 0 1 10 6 3 D53402 H12992 0 1 10 6 3 D61243 T61971 D61243 H77240	J. 0.10	H545906	0	-	2	3	-	103569	Human lymphocyte activation antigen 4F2 large subunit
T61971 T61971 D61243 N77240	2010	H12992	0	-	2	٥	3	D53402	Human fetal brain cDNA 5'-end GEN-108D03.
D61243 N77240 N77240						T	Γ	T61971	yb96f02.r1 Homo sapiens cDNA clone 79035 5'.
N77240						T		D61243	Human fetal brain cDNA 5'-end GEN-171G06.
1371131 0 0 10 1 2 T35761							T	N77240	yv44d02.r1 Homo sapiens cDNA clone 245571 5'.
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	H555168		187711	10401	700CECH	11232021	H610614	
	CTTC ACTC ACTTC	1/8 CATGOACTOR		- 12 CATCAAACCCCAA		180 CATCATCACCCCCC	(A)COCO ACACOCO CA	181 CATOGCCCACATCCCCA

Table 3 - Transcripts decreased in colon cancer

Transcripts decreased in only colon primary tumors compared to normal colon (51 genes)

NC: Normal Colon TU: Colon Primary Tumor CL: Colon Cancer Cell Line PT: Pancreatic Primary Tumor PC: Pancreatic Cancer Cell Line

Gene Name	ë.	letal gamma-actin.	atin 18		demandent professe (small subunit)	Human mKNA for calcium dependent process (Smeri	H.sapiens CpG island DNA genomic Miser Hagnicity of	zd30d02.r1 Soares fetal heart NoHH19 w Homo Sapicals	-end GEN-141D02.	4 N Q m (352) -:	Human thyroid hormone binding protein (1957) mixton.	cDNA clone 2/0343 3	zb06a05.r1 Soares fetal lung NbHL19W Homo sapiens	osuccinate synthetase.	Human mRNA for very-long-chain acyl-CoA dehydrogen	A, clone 173.	A. 3' end.	A 2 2 2 2 2 2 2 2 2 2 2 2 2 2 2 2 2 2 2	A Million of the Company of the Comp	profein	Marie Will (COX8) mRNa	Human cytochrome c oxidase subulint viti (Coxto) misting	Human Na, K-A I Pase appra-1 subunin mixaya, compress c	gb/R50350/R50350 yj59cu4.st Homo sapiens contractions	cDNA clone 133030 3.	e 3NHC0642.
	Human mRNA for beta-actin.	U.man mDNA for cytoskeletal gamma-actin.	11. DNA for extokeratin 18	Human mixix for cytoken	Human lipocorun 11 mr.	Human mKNA for calcium	H.sapiens CpG island DNA	zd30d02.rl Soares fetal hea	Human fetal brain cDNA 3-end GEN-141102	Unknown	Human thyroid hormone of	yy05d05.s1 Homo sapiens cDNA clone 2/0343.3	zb06a05.r1 Soares fetal lun	Human mRNA for argininosuccinate synthetase.	Human mRNA for very-lor	Human keratinocyte cDNA, clone 173.	Luman alpha-tuhulin mRNA 3' end.	Т	T	H.Sapiens Idl Illicato.	H. Sapiens mindy to bit process	Human cytochrome c oxid	Human Na, K-A I Pase alpn	gb R50350 R50350 yj59cu	yj59c04.r1 Homo sapiens cDNA clone 135030 3	Human Heart cDNA, clone 3NHC0642.
Accession	V00351	10000	X04096	X12883	D00017	X04106	265513	W61077	D60944		J02783	N33042	W07627	X01630	1743682	1720146	22.00	X00557	AA341633	X77956	X87949	104823	U16798	R50350	R50013	18000
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	Tag sequence	CATGGCTTTATTTGT	CATGCTAGCCTCACG	CATCOAAACCATCCA		4 CAIGCLICCAGGIIII		6 CATGGAIGACCCCC		8 CATGCGGACTCACTG	9 CAIGCCCCCCCCCAAA	10 CATGCCTGGAAGAGG	11 CATGGCC1GGCCA1C	12 CATGAGCAGGAGCAG	13 CATGAACGTGCAGGG	14 CATGGCCGCCCTGCA	15 CATGTGGGGAGAGGA	A CATGGCTGCCTTGA	TATE TO	18 CATGCGTTCCTGCGG	TO TOTOL A TOTOL OF	19 CATOTTOTT	20 CATOTA COTOTA TOG	ZI CATUTAGCICIATOS	22 CATGGTGCGCTAGGG	

				ł	ł	ł	+	9	Serrondas Homo saniens cDNA 5' end similar to ubiquinol
		73780311	80	•			26 TJ	T31329	cytochrome-c reductase, 6.4 kDa.
23		H694707	27	╁╌	12	3	61	n	Unknown
24	CATGCCTCCAGIAC	H388627	23	m	4	90	7 H		yr34d11.rl Homo sapiens CDNA clone 20/109 5 smith
22	CATGCCIGIGACAGC	H856806	24	2	∞	12	11	1	zd27c08.rl Soares tetal neart Notanis w tionic suprema
2 2	26 CATGLCACAGIGCCI	H49320	23	2	7	=			Human G I Pase (moc) may A, complete cos
27	CAIGAAIAAAGCIA	H1031929	23	2	13	15	_		Human mKNA tor cannounin, compact 223.
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2	32 CATGATGGCACGAG	H608326	2	_	9	_	\neg		ESTITION HOMO Sapteris CDIAS Our Similar
7	33 CA IGGCCAGACACC	HS15990	92	0	11	3			Human gene for apina 1 grouns.
3	34 CATGC11C11GCCCC	1102452	2	-	7	22	<u>х</u>		Human jun-B mKNA for Jun-B protein.
35	35 CATGACCCACGTCAG	1686458	× =	1	4	5			yj90e08.s1 Homo sapiens cDNA clone 130036 3.
36	36 CATGGGCTGCCTGCC	10001		+		-	R	R48449	yj67b10.s1 Homo sapiens cDNA clone 133787 3.
				†	\dagger	\dagger	12	Γ	yj72b03.s1 Homo sapiens cDNA clone 154253 3.
			!	1,	12	-	× ×		Human Na+,K+ ATPase gene exons 1 - 3 (alpha III is
5	CATGGAGGGCCGGTG	H567660	٩	,	:	,	┰		Tinknown
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3 ?	S CATO COLOR OF CALC	H153109	91	7	=	-			men and transport of the complete cds and tr
2)	Sy CAlgadecode	H774780	2	2	12	~	\neg		Homo sapiens por in (por) mixty, complete
3	CAIGGIICAGCIGIC	H181443	91	-	8	9	7		Juman /8 KDa gasu in-purioning process in the second secon
7	41 CATGCCTCGCTCAGI	0165901	15	1-	∞	6	0	U17077	Human BENE mRNA, partial cds.
7	42 CATGCAAATAAAGI	1202217	=	-	∞	0	3	U28369	Human semaphorin V mRNA, complete cus.
4	43 CATGTGCCGCCCGCA	0700701	15	c	9	4	3	D12038	Human HepG2 3'-directed Mool CLINA, clouic 31.30.
44	44 CATGGCAGTGGCCIC	11602137	14	-	-	5	<u> </u>	U77396	Human TNF-alpha inducible responsive element moves,
4	45 CATGCTGGGCCTGAA	H302137	-	-	0	2	г	229093	H.sapiens EDDR1 gene for receptor tyrosine Kinase.
7	46 CATGGCCCATIGGAG	COCCET	: :	c	6	2	0	T94990	ye38a04.s1 Homo sapiens cDNA clone 119962 3.
-	47 CATGAAGAAACCTC	H32192	1	·	1	1	Т	N69310	
L_			1		1	†		Γ	zb86e03.s1 Soares senescent fibroblasts NbHSF Homo sapiens cDNA
<u>L</u> .							_=	N98502	clone 310492 3'
			2	6	٧	6	4	F18838	H.sapiens EST sequence (007-X1-01) from skeletal m
4	48 CATGGAATGATTTCT	H>388/8	<u>*</u>	<u>, </u>	·		Т		zr21b10.s1 Stratagene NT2 neuronal precursor 937230 Homo sapiens
<u> </u>		CLC1C3H	12	0	3	3	8	AA226928	cDNA clone 664027 3'
4	49 CATGGCCTGGICCII	11717011	: =	0	-	-	0	M60047	Human heparin binding protein (HBPI /) IIINAA
8	SO CCATGGCCCACACAG	HOION		·					

51 CATGGGATTCCAGTT

2 W52456 zc45e09.rl Soares senescent fibroblasts NbHSF Homo

Transcripts decreased in both colon primary tumors and colon cancer cell lines compared to normal colon (130 genes)

NC: Normal Colon
TU: Colon Primary Tumor
CL: Colon Cancer Cell Line
PT: Pancreatic Primary Tumor
PC: Pancreatic Cancer Cell Line

			I	ŀ	۲	H		According	Gene Name
3	Tag Sequence	Tag Number	ပ Ž	TU	3		1	CCCSSION	Said Complements
•	O VIII O VIII O O II O	H382109	803	161	304	136 6	663	(12882	X12882 Human mKNA 101 cytoket attil 0.
_	CATGCCTCCAGCTAC	9003811	20%	282	402	142 4	497 F	F15636	H.sapiens mitochondrial EST sequence (UUZ112)
7	CATGCTAAGACTTCA	07600HH	3	┿	+	4	-		linknown
-	CATGGCCCAGGTCAC	H610997	3	2	7	+	4		11 inchardrial FST sequence (009-T1-21) f
<u>\</u>	CATGACCTTGGGA	H90022	512	348	93	43 2	ᆉ	F-1694U	H. Sapiens Illinocing and the Sapiens (FARP) mRNA
7	CATCACATTGGGTGA	H81583	504	92	4	0	┥	410050	M10050 Human liver ratty actu Dindring provent (creek)
	CATORCALIGORION	H622680	486	801	27	30	13 8	S61953	c-erbB3=receptor tyrosine kinase (ancinatively sp
0	CATGCCAAACCCTG	19885114	367	242	132	71 2	204	F15506	H.sapiens mitochondrial ES1 sequence (1-1-02) II our
	CATGAGCCLIACAAA	1646078	276	131	6	7	0	T39321	T39321 ya04c01.r2 Homo sapiens cDNA cione ou480 3.
∞	CATGGACCCAAGATA	0706461	2	1	+	╀	┞	124673	H24673 vl41a01.s1 Homo sapiens cDNA clone 160776 3'.
				1	†	\dagger	+		HIMGS02706 Human colon 3 directed Mbol cDNA, HUMGS02706,
					-			73888	225686 clone cm 1673.
					-	-	1	25.230	COURT ON A STATE OF 11 7105 7'
								T96160	T96160 ye09b02.st Homo sapiens curva cione 11/12/2
			13.0	ŝ	100	2	178	X64364	X64364 H. sapiens mRNA for M6 antigen.
0	CATGGCCGGGTGGGC	H617195	ŝ	8	+	- 1	4		Additional familian H chain mRNA complete cds.
15	O TOTTO GOSTATOS	H1026814	202	75	84	235	369	M11140	number of the complete of
1	CA10110000110000	H479577	201	120	0	=	m	L15203	Human secretory protein (F1.D) linking, compress cos.
	CA IGC I CCACCCAA (a) a)	0230031	ě	87	٧	3	61	X93036	X93036 H.sapiens mRNA for MAT8 protein.
12	CATGGCAGGGCTCA	HonnoH	2	3	,	╀	╁╴		vv07h09_r1 Homo sapiens cDNA clone 242081 5' similar to SP:A39484
L_		-	101	2	6	-07	39	H93844	A39484 ANDROGEN-WITHDRAWAL APOPTOSIS PROTEIN RVPI
=	13 CATGATCGTGGCGGG	C76477H			: =	╀	╀	F17001	H. sapiens mitochondrial EST sequence (011-11-13) f
7	14 CATGCAAGCATCCCC	H271574	3	Ŷ		╅	1	20000	Sonson Himan mRNA for keratin 19.
Ι <u>.,</u>	15 CATGGACATCAAGTC	H544012	189	5	•		<u> </u>	200	Shores 11 r.1 Soares fetal lune NbHL19W Homo sapiens cDNA clone
<u> </u>					_				301148 5' similar to gb: V00567 BETA-2-MICROGLOBULIN
		11000	178	- 2	7	340	139	W16632	
<u>=</u>	16 CATGGTTGTGTTAA	C1070/H			1	1	L		zo31h04.s1 Stratagene colon (#937204) Homo sapiens cDNA clone
								A 143804	A A 143804 588535 3'
_		_	_		_		_	TO CET U	

197 z192h02.s1 Stratagene colon (#937204) Homo sapiens cDNA clone	AA133597 512115 3	T53199 ya86c05.s1 Homo sapiens cDNA clone 68552 3'.	R00081 ye73c04.s1 Homo sapiens cDNA clone 123366 3.	M16364 Human creatine kinase-B mRNA, complete cos.	yf22e12.s1 Homo sapiens cDNA clone 12/030 3 similar to contains and	R09410 repetitive element	HUMGS0003915, Human Gene Signature, 3'-directed county		yq04h09.s1 Homo sapiens cDNA clone 196001 3 similar	2h78e12.s1 Soares fetal liver spleen INFLS SI Homo	W90374 CDNA clone 418222 3' similar to contains Alu repetitive element	X52003 H.sapiens pS2 protein gene.		_		1.	Ul4945 Intilial Mile Link chain mRNA complete cds.	M81457 Human calpactin 1 light chain mixty, compression CDNA sequence	C21047 HUMGS0002546, Human Gene Signature, 3 -uncond control of the Control of th	zo21h08.sl Stratagene colon (#93/204) Homo sapiens	Z FEC	zi68h06.s1 Stratagene colon (#937204) Homo sapiens cDNA	AA054072 clone 509819 3'	zo18g08.s1 Stratagene colon (#937204) Homo sapiens cDNA clone	AA132736 587294 3' similar to SW:LEG4 RAT P38552 GALECTIN-4	X04412 Human mRNA for plasma gelsolin.	X77658 H. sapiens mRNA for HLA-B*7301.	zo35c09.s1 Stratagene colon (#937204) Homo sapiens cDNA clone	AA146606 588880 3'	zo35g09.s1 Stratagene colon (#937204) Homo sapiens cLNA clone	AA146775 588928 3'	zo74g11.s1 Stratagene pancreas (#937208) Homo sapiens cuiva cione	AA161043 592676 3'	
-			0	9	-	4	-	_		 -	_	_	181	╁	╀	+	-		91		<u> </u>	+	_<	┝		1	2	╁	7 7	! -				
-		\dagger	0	<u>س</u>		2	-		-			26	╁	╁╾	:	= 8	-	156	24			T				2	2	;	4					
f		\dagger	-	92	T	4			t			-	, 4	; =	:	-	32	8	0			T		T		1	:	;	_					
r		\dagger	12	33	T	4			T	 T		۶	3 5			×	\$	8	37			1				-	7	3	7					
-		\dagger	174	172		163		_	1	1		15	3 5	3 9	3	2	149	145	126			T				133	27 [771	- 5					
-			H947654	H284132		H368200	200001					1110011	HOLLINGH	H350110	H1001401	H256186	H493039	H149715	H655433							10553011	H87//01	H930217	11667337	1001000				
				17 CTAGIGCICCIACC	18 CATGCACCCIGATO		19 CATGCCGCIGCACIC	-					20 CATGCTGGCCCTCGG	21 CATGCCCCCTGGATC	22 CATGTTCACTGTGAG	23 CATGATTGGAGTGCT	14 CATGCTGACCTGTGT	24 CAIGCIGACCIG:	25 CA LGAGCAGA CAGG	26 CATGGGAAAACAGAA							27 CATGTCACCGGTCAG	28 CATGTGCAGCACGAG		29 CATGGGAACTGTGAA				

ţ

					-	\mid	-	F	-197 mg et Stratagene colon (#937204) Homo sapiens cDNA clone
		· · · ·				_	AA	A A 088704 51 1239 3'	11239 31
		1100111	=	5	2	6	40 H	H00427 y	yi23g11.r1 Homo sapiens cDNA clone 149636 5'.
ä	30 CATGCGAGGGCCAG	1404117	=	;	╁	╫	+	Т	2063d03.s1 Stratagene pancreas (#937208) Homo sapiens cDNA clone
							AA	AA158715 591557 3'	91557 3'
T				<u> </u>			12	T08562 E	EST06454 Homo sapiens cDNA clone HIBBG31 3' end.
T				T	T	-		Ï	zm21a12.s1 Stratagene pancreas (#937208) Homo sapiens cDNA clone
		-					¥	78845 5	AA078845 526270 3'
];	C. TOT A A TITO A A A	H790417	=	9	-	0	X 0	3502 F	X73502 H. Sapiens mRNA for cytokeratin 20.
<u> </u>		H686762	=	38	48	45 4	43 J0	J03191 F	Human profilin mRNA, complete cds.
7		H761359	60	2	8	1 19) 	12629 F	U02629 Human smooth muscle myosin alkali light chain mKNA
		H758243	107	2	36	34 8	82 X(7059 F	X07059 Human M4-50 mRNA for HLA class I antigen.
<u>بر</u>		H1032614	5	=	4	3	37 FI	5592	F15592 H.sapiens mitochondrial EST sequence (001124) from
<u>:</u>	CATOLITAACOOCCO			T			_		zl74e07.sl Stratagene colon (#937204) Homo sapiens cDNA ctone
	0 × × 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0	H157779	106	17	7	~	6 AA	099880	AA053660 510372 3' similar to contains Alu repetitive element
۶	CATGCCCICCCAAG	/2//0011		1		\vdash	I	_	HUMGS04077 Human colon 3'directed Mbol cDNA, HUMGS04077,
							Ω	D25711	clone cm1210
			1		T	-	-		H.sapiens CpG DNA, clone 140c4, reverse read cpg14(Mitochondria
		27876	104	×	22	14	27 Z	Z56800 F	EST
37	CATGAGGTGGCAAUA	H204104	3 2	: =	0	╁	╁	M95174	Human guanylin mRNA, complete cds.
8	CAIGAIACICCACIC	1787087	ē	12	~	4	91		Unknown
23	39 CATGCTCGCGCTGGG	140420	2		+	╁	-		yn01b01.r1 Homo sapiens cDNA clone 167113 5' similar to SP:ZK783.1
		HK07514	2	32	28	37	65 R	R90863 (CE00760 ;
무	CATGGGGGCAGGCC	10000				-	┞	T24702	EST277 Homo sapiens cDNA clone 10H4.
_]:		H\$33666	8	3	42	82	87 ×	X95404	H.sapiens mRNA for non-muscle type cofilin.
7	CATOOCACCACACA	H338569	75	22	28	R	× 91	67325	X67325 H.sapiens p27 mRNA.
4		H70711	74	=	8	0.	31 F	16604	F16604 [H.sapiens mitochondrial EST sequence (009128) from
-	CATGACACACA						-		za16a03.s1 Homo sapiens cDNA clone 292684 3' similar to contains Alu
		H134304	69	59	_	3	z -	N69361	
44	CATGAGAATAGCTIG	20000					-		ze30b10.s1 Soares retina N2b4HR Homo sapiens cDNA clone
							¥	816510	AA015918 360475 3' similar to contains Alu repetitive element
									y114h01.s1 Homo sapiens cDNA clone 158257 3' similar to contains Alu
							<u> </u>	H26689	repetitive element; contains TARI repetitive element ;.
\bot						┢			2779h I .s. Soares NhHMPu SI Homo sapiens cDNA clone 681957 3
4	45 CATGCGCTGTGGGGT	H424875	89	٥	٥	~	23 A/	256365	AA256365 similar to WP:C33A12.7 CE03333

2c39e11.s1 Soares senescent fibroblasts NbHSF Homo sapiens cDNA	W47357 clone 324716 3'	zb90f03.s1 Soares senescent fibroblasts NbHSF Homo sapiens cDNA	W19276 clone 310877 3'	R07159 yf13h12.s1 Homo sapiens cDNA clone 126791 3'.	0 0 L02785	3 6 U11862	1 2 N93240 2b68b06.sl Homo sapiens cDNA clone 308723 3'.	NIB1986 Normalized infant brain, Bento Soares Homo sapiens cDNA	T16906 3*end.	yu22h07.s1 Homo sapiens cDNA clone 234589 3' similar to	H78256 SP:SBP_MOUSE P17563 SELENIUM-BINDING	EST47523 Homo sapiens cDNA 3' end similar to similar to Selenium-	T32362 binding protein,liver.	3 0 V00493 Human messenger RNA for alpha globin.	7 14	15 3	111 7	H03961	R33498 Jyh83f04.s1 Homo sapiens cDNA clone 136351 3'.	1	0 0 AA053043	1 30 F17394	21 8 Z13009	8 31 X15505	0 0 H14641	32 8 M20469		8 11 N50873	0 1 U79725	4	22 24	H52178 Jyt85h08.s1 Homo sapiens cDNA clone 23 1135 3.	T40539 ya05b02.s1 Homo sapiens cDNA clone 60555 3'.
_					0	0	-							0	7	2	2			_	0	5	=	5	0	=	_	~		-	14	_	
L		1			5	2	E			_				<u> -</u>	21	91	9		_	1	9	7	2	=	4	Ε	<u> </u>	5	-	2	11	_	-
		1		_	89	8	2					_		57	SS	2	2		1	_	2	S	\$	48	48	4		46	45	44	43	_	1
					H314109	H614731	97171H	2010						H344474	H550554	H87386	H236169	20.00			11862097	H723890	H977640	H650847	H929299	H686744		H800074	H545514	H673210	H41344		
					O ATTROCATA COLLA	46 CATGCATAGGTT	4/ CA100CCCACCAOO!	48 CA LOAGCICI ICOAG						TOOODAAOOOTAO	49 CATGOACGCGG	200000000000000000000000000000000000000	SI CATORCCCCCCCCCC	52 CATUATUCCOCAGAA				SA CATOGTA AGTGTACT	se CATOCITOCITO	\$\$ CATOCOTOTOC	ST CATGAGTGACAGA		200000000000000000000000000000000000000	59 CATGTAATCCCAGCA	60 CATGGACCAGTGGCT	61 CATGGGCACCGTGCT	62 CATGAAGGACCTTT	22	

					Ì	ł			
		l					¥	303091	AA303091 EST12940 Uterus tumor I Homo sapiens cDNA 3' end
						\vdash		7	za52d02.ri Soares fetal liver spleen INFLS Homo sapiens cDNA clone
;		H 599903	43	•	12	24	13 W	02429	W02429 [296163 S.
3	CATGCCAGCTCCTGT				T	_	ž	N20325	yx44c11.s1 Homo sapiens cDNA clone 264596 3.
						_	Ž	N45127	yz13c12.s1 Homo sapiens cDNA clone 282934 37.
				T	T	╁	-		zb38c11.s1 Soares parathyroid tumor NbHPA Homo sapiens cDNA
							Ž	N90407	clone 305876 3'.
	OTTO TOTOTOR OF TO	H972720	5	2	4	22	S U	U03106	Human wild-type p53 activated fragment-1 (WAF1) mK
હ	(4 CATGTGTGCTGGTTG	2777				\vdash			zc11f01.s1 Soares parathyroid tumor NbHPA Homo sapiens cDNA
,	A D D D D A A A D A D T A D	H65878	42	91	7	12	11 W	W37827	clone 322009 3'
S	CATUACAAACCCCCA				T		_		gblW15332lW15332 zc16d10.s1 Soares parathyroid tumor North A
								W15332	Homo sapiens cDNA clone 322483 3'
				T	T	╁		Ï	zc04g10.s1 Soares parathyroid tumor NbHPA Homo sapiens cDNA
						-	<u></u>	W32410	clone 321378 3'
				T	T	T	Z	N32312	yw82c01.s1 Homo sapiens cDNA clone 258720 3'.
		1000011	į	1	=	-	0	1151478	Human sodium/potassium-transporting ATPase beta-3
99		H828331	;	,	: -	╁	十	-	Unknown
67	CATGACTGTGGCGGC	H120019	;	·	+	+			2244fil el Stratagene muscle 937209 Homo sapiens cDNA clone
_			Ş	٦	-		24 AA	180815	A A 1808 5612333 3' similar to contains Alu repetitive element;
89	CATGGTAGCAGGTGT	H/3028/	3	1	=	╫	1		yh87e04.51 Homo sapiens cDNA clone 136734 3' similar to contains Alu
							~	R34696	repetitive element;.
				T	1	T	-	T	yh87e04,s1 Homo sapiens cDNA clone 136734 3' similar to contains Alu
							~	R34696	repetitive element;.
					T		-		zq06e03.s1 Stratagene muscle 937209 Homo sapiens cDNA clone
							<u>\{</u>	194497	AA 194497 628924 3' similar to contains Alu repetitive element
					T				hbc760 Homo sapiens cDNA clone hbc760 3'end similar to nonspacific
		H43508	40	12	0	۳	0	T11144	crossreacting antigen.
8	CATGAATCACAAAA	200001					1		zi67e01.s1 Stratagene colon (#937204) Homo sapiens cDNA clone
								1058357	AA058357 509688 3' similar to TR:G189087
\perp			$oldsymbol{\perp}$			T		C05803	similar to none
						T			zo31e02.s1 Stratagene colon (#937204) Homo sapiens cDNA clone
- 1		H167606	40	=	4	4	5	1143765	AA143765 S88506 3'
۲	70 CATGAGGATGGTCCC					-	-		zp45b09.s1 Stratagene HeLa cell s3 937216 Homo sapiens cDNA clone
							<u> </u>	1179299	AA179299 612377 3'

CATGCCAAAGCTATA		×		_	7		M35252 Human CO-029.	
OCCUPACION OF THE PROPERTY OF	H326306 H434907	<u>ه</u>	- ∞	9 0	╁	╀	R87448 ym89c10.s1 Homo sapiens cDNA clone 166098 3'	one 166098 3'.
CATGGGGGGGGGG	H618121	38	6	2			X79882 H.sapiens Irp mRNA.	
CATGCCCCGAAGCC	H349706	37	9	-	\dashv	+	_	complete cds
75 CATGATTTCAAGATG	H259108	37	-	0	\dashv	0 103037	Human carbonic annydrase II innyd, comprete con	i, complete cas.
76 CATGGCCCAGTGGCT	H611050	2	~	٠,	+	10	1	regulator mRNA
77 CATGATGGTGGGGA	H241323	%	7	ام	<u>.</u> ا	╁	Т	ine/threonine kinase
78 CATGCCTGCCCCCT	H386390	33	2	7	, - 	X00180	_	nolete sequence).
79 CTAGTGGAAAGTGAA	H950457	×	1	- -	<u>.</u>	┿	_1	te cds.
80 CATGGTCATCACCAC	H740629	34	-	\$	 	7.00		
CATGETTATGGTCCC	HS11670	34	-	0	<u>س</u>	1 AA287	AA287021 2557c03.s1 Soares NbHTGBC Homo sapiens cDNA clone 701572 3'	o sapiens cDNA clone 701572 3'
DIOTOGOTOGO	H502136	34	m	4	=	5 T55226		2007 1 II- 10 -
						R37446	yf56e10.s1 Homo sapiens cDNA clone 20129 3 similar to go. Act 173 INTER-ALPHA-TRYPSIN INHIBITOR COMPLEX COMPONENT II	TOR COMPLEX COMPONENT II
						AA400	AA406180 zu65c08.s1 Soares testis NHT Homo sapiens cDNA clone 742862 3'	o sapiens cDNA clone 742862 3'
	H610982	33	5	0	0	2 R09752	2 Unknown	13 673671
JUCCANOSCEC	111047673	33	7	0	4	2 R81530	0 yj02b10.rl Homo sapiens cDNA clone 14/34/3	one 14/34/ 3.
CAIGITITACIONI						T323	T32348 EST47211 Homo sapiens cDNA 3' end similar to None.	end similar to None.
						W57810		119W Homo sapiens cDNA clone
							zt47e12.s1 Soares ovary tumor NbHOT Homo sapiens cDNA clone	4OT Homo sapiens cDNA clone
						AA39	$\overline{}$	in the second second
SOLULIA SOLUTION SIL	H387054	32	2	-	9	32 X63187	\neg	H.sapiens HE4 mRNA for extracellular proteinase innibitor notification
CATOCATORAGE	H96931	32	9	4	8	9	Unknown	36222 31 similar to ab: M33087
200000000000000000000000000000000000000	03100011		_	٥	0	0 R46266		yg52g07.s1 Homo sapiens cDNA clone 30232.3 Similar to go.tv33737 CARBONIC ANHYDRASE I
CATGCCTTCAAAICA	H893564	: =	-	4	1	├	_	lone 261490 3'.
CATGICGGAGCIGII	2000	-					zo97h01.s1 Stratagene ovarian canc	zo97h01.s1 Stratagene ovarian cancer (#937219) Homo sapiens cUNA
						AA17		071064 21
		L				H99212	2 Jyx15g08.s1 Homo sapiens cDNA clone 201834 3	lone 201834 3.

					T	I	-		zk10e12 s1 Soares pregnant uterus NbHPU Homo sapiens cDNA clone
							_ ∢	AA029975 470158 3'	70158 3'
ç,	CATCOOL	H666539	8	ø	2	32	22	M75161	H.sapiens granulin mRNA, complete cds.
် ခြ	ON CATGETCCACTAACC	H1003970	30	7	3	\dashv	4		gblU53204(HSU53204 Human plectin (PLECT) mKNA, complete cos.
\$ 5	91 CATGTCTGGGGAT	H752297	56	-	~	م	F	T60135	yc22a06.s1 Homo sapiens cuive cione o 1334 5 .
								120402	gb U6/y63 H5U6/y63 Human lysophiosphonipase nomere (1.10 1.1)
					†	\dagger	\dagger	Т	vh39a12.rl Homo sapiens cDNA clone 132094 S' similar to gb:D26129
		H984414	29		0		-	R23595 F	RIBONUCLEASE PANCREATIC PRECURSOR (HUMAN)
76	CATOLINACCECTEC							\$77094	yj83c08.s1 Homo sapiens cDNA clone 155342 3' similar to gb:D26129 RIBONICI EASE PANCREATIC PRECURSOR (HUMAN);
			1	+	†	\dagger	\dagger	T	vi84h01 s.1 Homo sapiens cDNA clone 145969 3' similar to gb:D26129
								R79191	RIBONUCLEASE PANCREATIC PRECURSOR (HUMAN);
								N40965	yjS6c03.s1 Homo sapiens cDNA clone 152740 3' similar to gb:UZ6129 RIBONUCLEASE PANCREATIC PRECURSOR (HUMAN);.
				1	1	\dagger	+	Т	2.25k12 r.1 Soares overy filmor NHOT Homo sapiens cDNA clone
									755687 5' similar to TR:G459890 G459890 OVEREXPRESSED IN
		U221020	28	~	~	4	<u>v</u>	A410947	AA410947 TESTICULAR TUMORS
93	CATGATGACGCICAC	1431047	;	1	1		Т	H02520	yj40c11.r1 Homo sapiens cDNA clone 151220 5'.
						+	\vdash	t	zo12g08.r1 Stratagene colon (#937204) Homo sapiens cDNA clone
									586718 5' similar to TR:G459890 G459890 OVEREXPRESSED IN
							<u></u>	A 130551	AA130551 TESTICULAR TUMORS.

						\Box			zd33c10.s1 Soares fetal heart NbHH19W Homo sapiens CDNA clone
94	CATGCACCTGTCATC	H286420	28	~	9	~	4	W68230	34,243U 3' SIMILIAT TO CONTAINS AND TEPCHINE CONTAINS A IU
								R 89822	ypydauz.si nomo saprems como ciono recordo como recordo como repetitive element:
					T	\dagger	\dagger	Т	
									zk69e08.s1 Soares pregnant uterus NbHPU Homo sapiens cDNA clone
							_	1A053322	AA053322 488102 3' similar to contains element MEK6 repetitive element
80	CATGGATCCCAACTG	H578824	27	-	-	24	-	V00594	Human mRNA for metallothionein from cadmium-freated cells
	TOOOT	HS10123	27	_	٠	0	9	H43742	YPZ IQUS, EI HOIRO Sapiens CENTA CIONE TOCKES CONTRACTOR EZRIN
8	% CATCLIAGAGGGG	H238025	27	4	3	-	0		embly09616 HSICE H.sapiens mRNA for putative carboxylesterase
9	97 CATGATGGCCCATAC	P88105H	1/2	<u> </u>	, 0	7	0	V00497	V00497 Human messenger RNA for beta-globin.
8	98 CATGCCAAGAAGIG	ПЈ71001			.]		1		

on Ica TCTA COTOTICA TT	H810468	27	à	1	=	12 X	65614	X65614 [H.sapiens mRNA for calcium-binding protein S100P.
100 CATGIACCICATION OF TO A T	H233106	78	0	2	0	2		
200000000000000000000000000000000000000						-		emb Z69881 HSSERCA3M H.sapiens mRNA for adenosine
101 CATGITCIGIAGCCC	H1014566	25	5	0	4	0		triphosphatase, calcium
102 CATGCCTGTCCA	H388582	24	-	2	-	3 T9		ye65c02.r1 Homo sapiens cDNA clone 122594 5.
200000000000000000000000000000000000000					_	1	T87539	yd89f09.s1 Homo sapiens cDNA clone 115433 3'.
					\vdash	-		gblAA347726JAA347726 ESTS4132 Fetal heart 11 Homo sapiens cDNA
103 CATGTATGATGAGCA	H844682	23	4	0	_	0		S' end similar to transmembrane secretory component
104 CATGCTGGCAAAGGT	H500747	23	0	0	0	0		
105 CATGCTTGATTCCCA	H517078	23	4	4	11	7 1		Homo sapiens bone-derived growth factor (BPGF-1) m
106 CATGCTTGACATACC	H516402	22	0	0	7	7 X	X68277	phase
								Human N-benzoyi-L-tyrosyi-p-amino-benzoic acid
107 CATGGCTGGCACATT	H649492	22	5	0	-	2	_	alpha subunit (PPH alpha) mRNA, complete cds
108 CATGTCTGAATTATG	H909556	21	-	_	_	× -	X16354	Human mRNA for transmembrane carcinoembryonic antigen (CEA)
				-	_			H.sapiens mRNA for Gal-beta(1-3/1-4)GlcNAcalpha-2,3-
TOVOUV	H657554	21	_	_	 °	<u>×</u> ۳	X74570	sialyltransferase
109 CATGOGAAGAGACACT	20010011		1	T		╁╴	⇈	yo45d01.s1 Homo sapiens cDNA clone 180865 3' similar to contains
* OOGETOEO OE * O	H646998	20	2	•		0	R87768	PTR5 repetitive element
וומראומתרובוובבבבא	2000			1		\mid		yo36g07.s1 Homo sapiens cDNA clone 180060 3' similar to contains
							R85880	PTRS repetitive element
CATOTOTOTO	1114245	2	7	0	4	3	20826	L20826 Human I-plastin mRNA, complete cds.
TI CATOTA ATTACANT	H802708	0	1~	0	-	7	Z50751	HSB4BMR H.sapiens mRNA for B4B
וייי כאוסואאוווסכאוו							U77085	Human epithelial membrane protein (CL-20) mRNA, complete cds
							Y07909	HSPAPR H.sapiens mRNA for Progression Associated Protein
JJJJJJJJJJJTTTTTT	H764570	~	-	-	- - -	2	R48529	yj64g10.r1 Homo sapiens cDNA clone 153570 5'.
222222222222222222222222222222222222222						-		EST10a24 Clontech adult human fat cell library HL1108A Homo
* OTOTOOL * TTOL * O	H998127	17	0	0	_	0	T27534	sapiens cDNA clone 10a24.
THE CALCULATION AND A COLUMN	H663571	12	-	1~	4	0	T86124	yd84b04.s1 Homo sapiens cDNA clone 114895 3'.
- I S C A I C C C A C A C A C A C A C A C A C						╁╴	1-	zo15g05.s1 Stratagene colon (#937204) Homo sapiens cDNA clone
						<u> </u>	A 13 1008	AA131008 587000 3'
						-	R49945	yj58g11.s1 Homo sapiens cDNA clone 152996 3'.
							T57044	ya84h01.s1 Homo sapiens cDNA clone 68401 3'.
116 CATGCCAACACCAGC	H328787	17	-	0	0	0		
117 CATGAGGTGACTGGG	H178299	11	0	0	0	0		13 /20/31
118 CATGGCATGCA	H609654	9	0	0	0	0		gblR73013JR73013 yj94a09.rl Homo sapiens cDNA clone 126376 5.
110100								

		1	-	h	-	H	460012	14. 14. 14. man giraning nucleotide-binding regulatory protein
	H1039799	2	_	0	4	4	102013	Mulian Businia marian
CATOLOGICATION OF THE PROPERTY	H860776	5	-	-	_	0		
120 CATGICAGAGCOCIO			T			-		vv72h06.s1 Soares fetal liver spleen INFLS Homo sapiens
								CDNA clone 248315 3' similar to contains element PTR7 repetitive
	H1006014	4	_	•	•	7	N58523	element
121 CA1011CCCC011CC	H814011	4	-	0	0	0		Unknown
122 CATGIACGUIGIGG	710000	2	c	-	4	2		Unknown
123 CATGCTCAGAACT 1G	017//bH	-	,	+	+	╀	0730674	Uman carcino-mbryonic antigen mRNA (CEA), complete cds.
LACATGGGACTAAATGA	H662543	2	_		_	1	04C67IV	THINGSOATS A LINEAR COLOR 3 directed Mbol CDNA, HUMGS04154,
					_			The state of the s
TT & COCOTTO COT . C	H653988	12	0	0	0	_	D25786	D25786 clone cm0215.
125 CA100C11000A11			1		r	-		vc36e02.rl Homo sapiens cDNA clone 82//8 5 Similar to go. Lo. 705
							T73613	LIVER CARBOXYLESTERASE PRECURSOR
		!	1,	1	6	-		1 lakaowa
LISECATGACCCAACTGCC	H86138	12	9	5	-	+		Simple of the state of the services of the 120220 3.
000100 + 01001 071	H491894	12	0	0	7	7		[gb] 195615[195615] yequeus.st monto sapiens ectivities
127 CATGCTOAACCTCCC			T			-		zr19611.s1 Stratagene NT2 neuronal precursor 937230 Homo sapiens
	0112011	=	_	_	2	0	A226797	AA226797 cDNA clone 663837 3'
128 CATGCAAGAGTITCI	7011/71		,	+	1	+		zq97h01.s1 Stratagene NT2 neuronal precursor 937230 Homo sapiens
							A218730	AA218730 CDNA clone 649969 3'
			1	†	1	+		vn57f10 r1 Homo sapiens cDNA clone 191563 5' similar to gb:M90657
		:			•		H38178	
129 CATGGTCCGAGTGCA	H743610	=	- 1	,	,	\dagger		
LINDICATIONTICAC	H1043445	=	0	0	7	3		Olivilowii

cell lines compared to normal colon (78 genes) Transcripts decreased in only colon cancer

NC: Normal Colon
TU: Colon Primary Tumor
CL: Colon Cancer Cell Line
PT: Pancreatic Primary Tumor
PC: Pancreatic Cancer Cell Line

TILL PT Accession	moer 110 755 411 161 333 F15516	173 F12396	227 603 566 138 249 173 1125	704 452 595 235 80 314 LUG441	566 444 357 114 64 191 F15553	132 385 402 223 278 132 XS1525	360 446 171 76 161 F16402	262 202 527 78 14 83 U09500	22 200 169 98 17 223 F15744	194 127 70 21 75 F15511	147 192 04 49 57 F18587	14/ 103 01 CO A7 H03983	145 160 91 09 47	5322 124 194 63 111 51 A/4301	595 98 106 17 183 10/ M1//33	616 97 186 17 41 49 040913	638 67 48 25 75 34 XU36U7	3465 64 121 28 24 15 D54113	5339 60 33 17 13 15 X14758	56 41 4 31 3	434 53 271 6 30 5 D50954	436 49 35 10 100 36 M11233	3719 49 37 21 27 15 U25801	45 26 18 23 15 U31215	44 56 2 6 1 S79597	42 32 15 20 5 T48809	
-	╀	+	4	┥	_	⊢	╀	╀	-	╀	+	+	+	4	-	-			_	-		H	├	H	-	-	
⊦	╁	╁	┥	_	-	├	╁	╁	+	╁	┿	+	\dashv	-	\dashv	\dashv		\vdash	-	┝	┝	-	┝	╁	╀	╀╌	
N box	╅	2	H260227	H933704	H1002566	H1115412	7907111	11001082	111777	7/7/H	H4/8249	H885354	H103075	H1025322	H1027595	H214616	H941638	H136465	H196339	H656389	H965434	H527436	H763719	H765509	H704160	H763567	
	T	1 CATGCACCTAATTGG	CATGATTTGAGAAGC	Т	Т	- 1	s CATGCCACIGCACIC	-1	7 CATGCACIACICACE	8 CATGAAAACATICIC	9 CATGCTCATAAGGAA	10 CATGTCGAAGCCCCC	II CATGACGCAGGGAGA	12 CATGTTGGCCAGGCT			┰	_1.								23 CATGGGGIIGGGIIG	つう うりつうつう こうつ マンニャン

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									2080f04.s1 Stratagene ovarian cancer (#937219) Homo sapiens
								AA165679	AA 165679 cDNA clone 593215 3'
		11028404	,	7	-	-	4	zv40a02.s AA411012 756074 3'	zv40a02.s1 Soares ovary tumor NbHOT Homo sapiens cDNA clone 756074 3'
4	CAIGIAIAGICCICI	1100001		T	1				zl92g08.s1 Stratagene colon (#937204) Homo sapiens cDNA clone
								AA133595 512126 3'	512126 3'
						Γ			zt56b12.s1 Soares ovary tumor NbHOT Homo sapiens cDNA clone
								AA292774 726335 3	726335 3'
45	CATGGGTCCTCTT	H710520	20	7	2	2	2		yj73h02.rl Homo sapiens cDNA clone 154419 5' simil
49	CATGATGGGCTTGAT	H240121	19	4	0	~	3	D20113	Human HL60 3'directed Mbol cDNA, HUMGS01086, clone
47	CATGCTGCCCCCAT	H496981	61	5	0	_	4	\neg	Unknown
48	CATGITCICTACACA	H1013522	61	4	-	8	2		Human TSC-22 protein mRNA, complete cds.
8	CATGAAGAAGCAGGG	H33355	82	4	2	2	8	. [yj05g03.r1 Homo sapiens cDNA clone 147892 5'.
: 5	CATGAGTAGGTGGCC	H183018	82	131	2	17	7	D51021	Human fetal brain cDNA 3'-end GEN-007D07.
∤ 5	CATGACAGTGTGTGT	H77551	81	5	3	0	8	D26146	Human DNA for putative protein kinase.
: s	CATGGGAAAAGTGGT	H655547	81	13	3	70	1	M11465	Human alpha-1-antitrypsin mRNA, complete cds.
3 2	CATGAAGAAGCTC	H32926	17	4	0	~	-	R78188	yi81g01.r1 Homo sapiens cDNA clone 145680 5!
3	CATGACACCCATCAC	H70965	17	4	0	0	0	M22406	Human intestinal mucin mRNA, partial cds, clone SM
5 ž	CATGAGATCCCAAGG	H144707	17	81	0	0	0	T24507	EST082 Homo sapiens cDNA clone 3E6
5	200000000000000000000000000000000000000								za63a11.s1 Homo sapiens cDNA clone 297212 3' similar to
								N79237	PIR:S49589 S49589 cortical granule lectin - African clawed frog ;.
								T31354	EST30893 Homo sapiens cDNA 5' end similar to None
3	CATGAATAGTTTCCC	H52214	91	4	0	0	0	H54696	yq92e02.s1 Homo sapiens cDNA clone 203258 3' simil
2 5	CATGORGABAGGATG	H295060	91	6	0	0	0	M22430	Human RASF-A PLA2 mRNA, complete cds.
÷. =	CATGCATAGCATTG	H654976	91	4	~	∞	_	AA374631	EST86866 HSC172 cells I Homo sapiens cDNA 5' end
2.	2000000	21.							zn93g08.r1 Stratagene lung carcinoma 937218 Homo sapiens
								AA137163	AA137163 cDNA clone 565790 5'
									zk10f05.s1 Soares pregnant uterus NbHPU Homo sapiens cDNA
								AA029320	AA029320 clone 470145 3'
9	A TET OF OF THEA	H948543	15	2	0	-	0	D25681	Human colon 3'directed Mbol cDNA, HUMGS04047, clon
									zr72g02.s1 Soares NhHMPu S1 Homo sapiens cDNA clone 668978
								AA253331	3,
								H05110	yl75f07.s1 Homo sapiens cDNA clone 43778 3'.
Ş	CATGCCATCGTCGTT	H341720	15	8	-	-	10		Unknown
3 2	$\overline{}$	H529013	14	23	0	0	0	AA297150	AA297150 EST112734 Colon I Homo sapiens cDNA 5' end
	П						İ		

zd42c12.s1 Soares fetal heart NbHH19W Homo sapiens cDNA clone W68073 343318 3' similar to contains Alu repetitive element; 0 0 = H874226 78 CATGTCCCCGTTACA

Table 4 - Transcripts increased in pancreas cancer.

SAGE Tags elevated only in Pancreatic Tumor

No Normal Colon

Tu Colon Tumor

CC Colon Cancer Cell Line

PT Pancreatic Tumor

PC Pancreatic Cell Line		Ļ			James Manne
The Commence	Tap Number NC Tu	Tu CC PT PC		Ē	Cent Ivalia
1 ag Schming	10	1 3	11 Examples R38305		yh95b04.s1 Homo sapiens cDNA clone 13/455 3
CATGAAGCAAACCA	1				2k95b03.s1 Soares pregnant uterus NbHPU Homo sapiens CUNA clone
				AA126719	490541 3'
				Г	zk51c03.s1 Soares pregnant uterus NbHPU Homo sapiens cDNA clone
				AA044296	486340 3'
				Г	2133c08.s1 Soares pregnant uterus NbHPU Homo sapiens cDNA clone
				AA131586	503726 3'
					zo71h12.s1 Stratagene pancreas (#937208) Homo sapiens cDNA clone
	H9408	5 2 21	3 Examples	Examples AA157983	592391 3'
2 CATGAAAGCAGTTTA	1	1			zt54e04.s1 Soares ovary tumor NbHOT Homo sapiens CDINA Clone 1201174
				AA292929	31
					zo78c07.s1 Stratagene pancreas (#937208) Homo zo78c07.s1 Stratagene
				AA159306	pancreas (#937208) Homo
					vi70h01.s1 Homo sapiens cDNA clone 154129 3'
		+			vb99f08.s1 Homo sapiens cDNA clone 79335 3'
	1	1	13 Byamples X52426		H. sapiens mRNA for cytokeratin 13
3 CATGAAAGCGGGGCT	5	5	1	V51608	H caniens snasmolytic polypeptide (SP) mRNA.
4 CATGAAATCCTGGGT	H13803 0	1 10	1	20107	11.34 1.1 1. 1. 1. 1. 1. 1. 1. 1. 1. 1. 1. 1.
S A A D A D D T A A D T A D A D	H14865 0	0 1 0	13 Examples N70419	N70419	2861012.51 Home Saprems Court at the Court of the Court o
בעו פעעי פעעי				AA411599	zv16g01.r1 Soares NhHMPu S1 Homo sapiens CUNA Clone 733840.3
				4 4410508	zy 16g01 s1 Soares NhHMPu S1 Homo sapiens cDNA clone 753840 3'
		<u> </u>		2000	2186212.81 Stratagene colon (#937204) Homo sapiens cDNA clone 511558
	1 1247171	8	13 Examples	Examples AA115723	31
6 CATGAACCAGTTTG1	1]_		zo19e04.s1 Stratagene colon (#937204) Homo sapiens cDNA clone 38/338
				AA132875	3'
					2044a06.s1 Stratagene endothelial cell 937223 Homo sapiens CLINA Clone
				AA147677	589714 3'

					t			2081h12 s1 Stratagene hNT neuron (#937233) Homo sapiens cDNA clone
	-						AA206883	648071 3'
	H30689	<u>-</u>	7 13	12	=	Examples R51318		yg72f03.s1 Homo sapiens cDNA clone 38681 3'
CATGAACICIIGAAG		1						EST82235 Homo sapiens cDNA 3' end similar to None
						,	AA412071	2165h12.s1 Soares testis NHT Homo sapiens cDNA clone 727271 3'
K K COURT OF THE STATE OF THE S	H31221	+	8 9	७	200	Examples N63154		yz37f12.s1 Homo sapiens cDNA clone 285263 3'
S CATGAACTGCTTCAA	1221 (11		L					yc81h04.s1 Homo sapiens cDNA clone 22603 3'
		╁	_		T			2146f04.s1 Soares pregnant uterus NbHPU Homo sapiens cDINA cione 2049
		+	-				AA045773	zl68b12.s1 Stratagene colon (#937204) Homo sapiens
E WOOD DEBUT A KOMAN	H32405	-	0	∞	E	Examples X07819		Human pump-1 mRNA homolog. to metalloprotemase,
y CAI GAACT I GGCCAI		+	1				L22523	Human matrilysin gene, exon 5
COCCOMPANAMANA	H36183	2	10 14	12	23	Examples R72650	272650	yj95e05.s1 Homo sapiens cDNA clone 156512 3'
		-						Second Second Second No. 19 No. 19 No. 19 Second Se
								2030602.31 Suates total liverity of the PCOLI P16654 PERIPLASMIC
					•		1000 K	144238 3 SIMILIA IO 34 COLLA COLLA COLLA COLLA CALLON TOLERANCE PROTEIN CUTA
		-	4		1		10701	vio 5-015 s.1 Homo sapiens cDNA clone 156512 3' similar to
								SP-CYCY ECOLI P36654 C-TYPE CYTOCHROME BIOGENESIS
							R72650	PROTEIN CYCY
		+	+		T			
								zp61a11.s1 Stratagene endothelial cell 937223 Homo saptens CDNA ctone
								624668 3' similar to SW:CUTA_ECOLI P36654 PERPLASIMIC
							AA181976	DIVALENT CATION I OLERANCE HIGH AG for the Lok SH2 domain
				- ;		146751	1146751	Human phospholytosine independent again, per as more commented mRNA, complete cds
II CATGAAGGGAGGGTC	H43180	عاه	2 0	1	15	Examples J03077	103077	Human co-beta glucosidase (proactivator) mRNA
CATGAAGTTGCTATT	H40/20	1		1	•		M86181	Human prosaposin (PSAP) gene
		+	+				D00422	Human sphingolipid activator proteins, mRNA
		+	+				103015	Homo sapiens sphingolipid activator protein 1 mRNA
		\dagger	+				M60255	Human mutant cerebroside sulfate activator protein
	H57345	10	-	5 2	2	No Match		
CALGAAIGAAAAAA	H66031	=	4 24	5	8	Examples N22375	N22375	yw37d01.s1 Homo sapiens cDNA clone 254401 3
14 CATGACAAACIGIGG		+						2n20e01.s1 Stratagene neuroepithelium N12KAMI 93/234 Homo sapiens
							AA084643	cDNA clone 547992 3'
		$\frac{1}{2}$	$\frac{1}{2}$]]			

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AA046253 376682 3' H67396 2 7 7 16 37 Examples Z58016 H.sapiens CpG DNA, clone 26c7,	AA151668	H71151 0 1 0 2 14 Examples AA1556464	AA025673	H85924 0 8 5 13 4 Examples X02491	104164 Human interferon-inducing process of the membra vegos H saniens mRNA for interferon-induced 17kDa membra	12 7 Evamples X56841	H90050 1 4 2 13 7 X64879	H91579 49 22 45 70 94 Examples M21186	M61107	H97158 0 3 0 28 17 Examples D00244	K02286	M15476 Human pro-urokinase mkNA, complete cds	H103912 0 1 0 11 2 Examples L08835	M87313	CATG H113380 2 4 4 5 20 Examples H44451 yo75806.s1 Homo sapiens cDNA clone	AA157329 KD PROTEIN	2c32g06.s1 Soares senescent fibroblasts NbHSF Homo sapiens CDNA clone	W46455 KD PROTEIN
S CATCACTOACTA			In CATGACACCCIGIGS	TT CATCACTO TTGGATT			CATGACCCTTTAACA		IN CAT GACCGCCG1661	ないながらからなってなっている	שררו פו פערכע			פארפרררופרוב	22 CATCACGTGTGATG			

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		2	,	15	-	Examples M92357		Homo sapiens B94 protein mRNA, complete cds.
23 CATGACTCAGCCCGG	H119383	5		1				
	1123421	- 6	-	53	22	Examples X64875		H. sapiens mRNA for insulin-like growth factor binding protein 3
24 CATGACTGAGGAAAG	17667111	,	<u> </u>	Ŀ				Human growth hormone-dependent insulin-like growin jactor officialing
							M31159	protein 3
		+	\dagger	-	-		M35878	Human insulin-like growth factor-binding protein-3
			╁	_				insulin-like growth factor binding protein 3 (3) region?
	A)CACIU	-	10		6	Examples U65932		Human extracellular matrix protein I (ECMI) mkNA
25 CATGACTGCCCGCTG	חוקבקום	1	1	_	_			Human extracellular matrix protein 1 (ECM1) gene, exon 9
		1	t	\vdash	\vdash			zo03f09.s1 Stratagene colon (#937204) Homo sapiens cLivia cione colon
	80696171	4	0	7	72	Examples	Examples AA148916	31 SNIA Juna 586657
26 CATGACTGTATTTC	D120200		+		L		Г	zo12a11.s1 Stratagene colon (#937204) Homo sapiens culva cione Journal
							AA129137	3'
			\dagger	ig				2185g09.s1 Stratagene colon (#937204) Homo sapiens civing cloud of 1720
							AA115437	31
			\dagger	\downarrow	\downarrow		Π	zi87e07.s1 Stratagene colon (#937204) Homo sapiens cDNA clone 211020
							AA126967	3'
		1	1	+		E.omalec D24613		vh36c03.rl Homo sapiens cDNA clone 131812
27 CATGAGCACTGCAGC	H149395	1 2		<u>-</u>	9	CXallipics		mosens of Home saniens cDNA clone 186560 5'
JO DATE DE CAGGAGGGT	H150055	0	0	0	2	Examples H43243		Transport of Charles Indian Charles Indian Charles
	H162622	0 2	0	_	11	Examples X54942	1	H. Sapiells Chaise integrant integris NhHPi I Homo sapiens CDNA clone
יייייייייייייייייייייייייייייייייייייי					_	,		ZKOUBU (.St. Suates pregnam mores rose o recommendation of
n: newseartgacccc	H167446	1 7	12	의	피	Examples	Examples AA044081	480300 3
								486300 5' similar to PIR: A40533 A40533 cAMP-dependent protein kinase
							AA044211	major membrane substrate
	000000	,	6	18	7	Examples X14787	X14787	Class A, Human mRNA for thrombospondin.
11 CATGAGGTCTTCAAT	H1/0127		1_	-	=	Examples R27738	R27738	yh64f11.s1 Homo sapiens cDNA clone 134541 3'
12 CATGAGGTGCGGGG	H1/8002			╀	-			yj22f12.s1 Homo sapiens cDNA clone 149519 3' similar to 3F.2No37.3
							H00276	CE00436 ARSA
		-	L	-	-			zm19d07.s1 Stratagene pancreas (#93/208) Homo sapiens CD14A Cloud
	H181787		3	15	73	Examples	Examples AA076235	526093 3'
33 CATGAGTATCT GGGA	10000	L		+	-		H13159	yj16c04.s1 Homo sapiens cDNA clone 148902 3
		+		+	\vdash			2071e11.s1 Stratagene pancreas (#937208) Homo sapiens civing civing
							AA146632	592364 3'
	07770011	-	6	<u>~</u>	6	Examples X80062	X80062	H.sapiens SA mRNA.
34 CATGATACTTTAATT	D*/*07H	+	1_	+	+		U01691	Human annexin V (ANX5) gene
]	1	$\frac{1}{2}$			

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		-	+			ľ	VIDASA	Himan mRNA for vascular anticoagulant
		+	+	1	+		T	The state of the s
							M18366	Human piacental anticoaguiant protein (PAP) intenA
							M21731	Human lipocortin-V mRNA, complete cds
		\vdash					103745	Human endonexin II mRNA, complete cds
		\vdash	-		T			GAMMA-INTERFERON-INDUCIBLE PROTEIN IP-30 PRECURSOR
	H213518	7		5 25	_	Examples 103909		(HUMAN)
		\dagger	-					EST97384 Thymus II Homo sapiens cDNA 3' end similar to interferon,
	•						1	gamma transducer 1
TOTOS ACTOR OF ACTOR	H213679	2	9 25	22	156	Examples U09953		Human ribosomal protein L9 mRNA
10.000000000000000000000000000000000000		\dagger	_	1				Human ribosomal protein L9 mRNA, complete cds
		\dagger	╀	1				
			. <u>-</u>				D14531	Human mRNA for human homologue of rat ribosomal protein
		\dagger	+					zm03a05.s1 Stratagene corneal stroma (#937222) Homo sapiens cDNA
CATGATCAAGTTCGA	H213751	0	2	8	10	Examples AA063259		clone 513008 3'
	H210750	7	7 14	12	40	Examples L42856		RNA polymerase II transcription factor SIII p18 subunit mRNA
10 CATIGATIC GGCGCCA	H229502	╬		1.	₹	Examples Z59242		H.sapiens CpG DNA, clone 13a10, reverse read cpg1
3) CATENTERWALLING		+						
		\dagger	-		T			
JU CATGATGCGAAAGGC	H235531	7	3 12	3	22	Examples Z25820		H.sapiens mRNA for mitochondrial dodecenoyl-CoA dehydrogenase
		T	_				L24774	Homo sapiens delta3, delta2-CoA-isomerase mRNA
11 CATGATGTCTTCGTT	H243676	0	0	0	14	Examples M84711		40S RIBOSOMAL PROTEIN S3A (HUMAN)
1) CATGATGTCTTTTCT	H243710	-	7	<u>=</u>	7	Examples M62403		Human insulin-like growth factor binding protein 4
		\vdash	<u> </u>					Human insulin-like growth factor binding protein-4 (IGFBP4) gene,
-							U20982	promoter and complete cds
13 CATGATGTGTAACGA	H244487	6	4	5 44	8	Examples Z33457		H.sapiens mts1 gene.
			-				M80563	Human CAPL protein mRNA, complete cds
CO A A REGIONAL AND COMPANY OF THE PERSON OF	H270083	-	-	2	~	Examples N23207		yx70b09.s1 Homo sapiens cDNA clone 267065 3' similar to gb:L12330 THROMBOSPONDIN 2 PRECURSOR (HUMAN)
1000011000101			<u> </u>					1) (1) Spare over timor NHOT Home caniens cDNA clone 714188
	H286424		4	10	-	Examples	Examples AA285023	3' similar to gb:M33680 CD81 ANTIGEN (HUMAN)
יין כאופראכרופובכוו		†	1_	.!			Г	CD81 antigen
a a a d d d d d d d d d d d d d d d d d	H791889	10	6	2 3	161	Examples D78203		Neurosin
In CAT GCACT CAM LANA	(001/711		L	L				protease M
		1	$\frac{1}{2}$		1			

					t			
	H300971	- 6	0	- 0	10	Examples AA149942		2068d04.s1 Stratagene pancreas (#937208) Homo sapiens cDNA clone 592039 3' similar to TR:E218488 E218488 TRYPTASE
200000000000000000000000000000000000000								zp66b09.r1 Stratagene endothelial cell 937223 Homo sapiens cDNA clone 625145 5' similar to gb:M16937 HOMEOBOX PROTEIN HOX-B7
48 CATGCAGCGCGCCT	H301462	4	11 12	2	21	Examples	Examples AA187553 M16937	(HUMAN); contains element MER22 repetitive element HOX-B7
正しい正じ出出しいないのはない。	H307126	-	0	0	10	No Match		Alva and the
V CAT GCAGGI TGI CCI	H309109	7	9	77	11	Examples U14972	U14972	Human ribosomal protein 510 medya
SUCATOCAGICIONA CONTRACONACIONACIONACIONACIONACIONACIONACIONA	H316857	0	3	<u>س</u>	13	Examples U27293	U27293	Human leukotriene A4 hydrolase gene
OF CALGCATCCCG GAC		\vdash	-		П		103459	Human leukotnene A-4 nyarotase mutya, compress cus
			H				J02959	Human leukotriene A-4 hydrolase lingara, compress company of the manning of the m
TECOTOTATOTATOTA	H325080	0	7	5 13	~	Examples X82434	X82434	H. Sapiens mixton for cincing (ACR55) mRNA
Sales and Control of the Control of	H333138	6	7 17	7 18	7	Examples M88338	M88338	Human Serum consultation process (1922-202) and the consultation of mRNA
SOCOSOLO ACCORDANCE OF STREET	H339606	23	11 37	7 22	8	Examples U14971	014971	Human mossuma protein by medical mRNA
できたいのではないのです。	H344031	0	7	1 9	01	Examples L01697	L01697	Homo Sapiens atplia-1 type A v Couragen in Co.
SS CALGCCALLITICAGE	H344691	2		8 18	44	Examples X54079	X54079	Human mkNA for heat shock protein that any and
So CAT GCCCAAGCTAGC		\vdash	-				Z23090	H. sapiens mRNA for 28 KDa near shock protein
		-	-				X16477	Human mRNA tragment for estrogen-regulated 24h protein
		\dagger	+				S74571	estrogen receptor-related protein=27-kda heat snock protein
	00727	┸	15 13	10	15	Examples X69392	X69392	H. sapiens mRNA for ribosomal protein L26.
57 CATGCCCATCCGAAA	H34/489	3			3		1.07287	Human ribosomal protein L26 (RPL26) gene
	00005	-	+	7-	15	Examples U40434	U40434	Human mesothelin or CAK1 antigen precursor mRNA
SN CATGCCCCCTGCAGA	HSONOSA	+	\perp		1			Human mRNA for pre-pro-megakaryocyte potentiating factor, complete
						-	D49441	cds.
	10753611	1		o c	E	Examples U12819	U12819	Human p16-INK4 (p16) gene
SO CATGCCCGCATAGAT	H353401	>		1	1		1138945	Human hypothetical 18.1 kDa protein (CDKN2A) mRNA
		+	+	1				MTS1=multiple tumor suppressor 1/cyclin-dependent kinase 4 inhibitor
		_					\$69804	p16
		1	+	1			\$69822	CDK41=cyclin-dependent kinase 4 inhibitor
			+	\downarrow				tumor suppressor gene, P16/MTS1/CDKN2=cell cycle cycle negative
	•						S78535	regulator beta form
	7,87,5 ELI	~	-	5 14	34	Examples 247319	247319	H. sapiens mRNA for expressed sequence tag (clone 21fi7119)
60 CATGCCCTCCTGGGG	100/001		1	1		1		

							AA398406	2160h12.s1 Soares testis NHT Homo sapiens cDNA clone 726791 3'
	11370034	10	1	14	61	Examples U21049		Human DD96 mRNA
ol cargeegecerace	H3 /0034		+	: ?		Examples X03212		KERATIN, TYPE II CYTOSKELETAL 7
62 CATGCCTGGTCCCAA	H387925	" 	1	3	L			zp73f01.s1 Stratagene HeLa cell s3 937216 Homo sapiens cDNA clone
							A A 187637	625849 3'
		+	+	$\frac{1}{1}$	+		Т	2035g11.s1 Stratagene muscle 937209 Homo sapiens cDNA clone 611492
			,	,	- 2	Examples AA176457		3' similar to TR:G663269 G663269 BOLA
63 CATGCCTTTGAACAG	H392709	^	L	1			Г	zp35e11.s1 Stratagene muscle 937209 Homo sapiens cDNA clone 611468
							AA176541	3' similar to TR:G663269 G663269 BOLA.
		31	745	15	-	Examples X02492		Human interferon-inducible mRNA fragment
64 CATGCGCCGACGATG	H413644	1		2 6	\perp	Examples T53402		ya88g05.s1 Homo sapiens cDNA clone 68792 3'
65 CATGCTCAACAGCAA	C7+C/+H	1		╀				
								2d47g08.s1 Soares fetal heart NbHH19W Homo sapiens cDNA clone
							W69493	343838 3' similar to PIR:S24168 S24168 hypothetical protein - numan
	027	-	1,	15	-	Examples X13916	X13916	Human mRNA for LDL-receptor related protein
66 CATGCTCAACCCCC	H4/54/8	- 6		3 0	<u> </u>	Examples X80335	X80335	H.sapiens (24) Ferritin H pseudogene.
67 CATGCTGAGAAACTG	H4935/6	1		• •	2 2	Evamples X04878	X04878	Human mRNA for G(i) protein alpha-subunit
68 CATGCTGAGTCTCCC	H494454	=	Ł	7 2	2 :	Sydmin L	1114966	Human ribosomal protein L5 mRNA
69 CATGCTGCTATACGA	H498887	9	7		#	Examples Office	Tonkks	vd41 of 8.1 Homo sapiens cDNA clone 110846 3'
70 CATGCTGCTGAGTGA	H499247	╣	4	카	2	CAMILIPIES	200071	EST43791 Fetal brain I Homo sapiens cDNA 3' end similar to steroid
							A A 338799	hormone receptor hERR1
		+	1	+	╀		H97236	yv98b06.s1 Homo sapiens cDNA clone 250739 3'
		+		+	15	Examples C14084	C14084	Human fetal brain cDNA 3'-end GEN-018D10
71 CATGCTGGCGCCGAT	H501337	- 1	7 7 F		2 2	Examples D00017	D00017	Human lipocortin II mRNA
72 CATGCTTCCAGCTAA	H313161	5 6			-	Examples Z19574	Z19574	H. sapiens gene for cytokeratin 17.
73 CATGCTTCCTTGCCT	770+ICH	1	+	+	+		X62571	H. sapiens mRNA for keratin-related protein
		+	1	\dagger	+		X05803	Human radiated keratinocyte mRNA 266
	90100311	+	-	19	4	Examples X79067	X79067	H.sapiens ERF-1 mRNA 3' end.
74 CATGCTTTCTTCCT	06177CH		֓֓֞֜֓֓֓֓֓֓֓֓֓֓֓֓֓֓֓֓֓֓֓֓֓֓֓֓֓֓֓֓֓֓֓֓֓֓	15	12	Examples X51779	X51779	Human mRNA containing an Alu repeat
75 CATGGAAAAAAAAA	69747CH	_L	ł	1	+		X82240	H.sapiens mRNA for Toell leukemia/lymphoma I
	11575348	╅	4		122	Examples V00572	V00572	Human mRNA encoding phosphoglycerate kinase.
76 CATGGAAACAAGATG	OFCC2CH	+		+	-		D29018	Human keratinocyte cDNA, clone 001
		+	F	╁	╁		L00160	Human phosphoglycerate kinase (pgk) mRNA
	JE277416	100	35 10	001	38	Examples X05344	X05344	Human mRNA for cathepsin D
77 CATGGAAATACAGTT	117217	=						

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		-	F	-	 -		M11233	Human cathepsin D mRNA, complete cds
	01011010	1	2	4	92	Examples T90296		yd42f03.s1 Homo sapiens cDNA clone 110909 5 similai to or .N.151.7 CE00827
N CATGGAAATGATGAG	(27/20)	-					142	EST23523 Adipose tissue, brown Homo sapiens cDNA 3' end
		╁,		+,	ę	Byomplec A A 181811		zp64f07.s1 Stratagene endothelial cell 937223 Homo sapiens cDNA clone 624997 3'
CATGGAAGATGTGTG	H533436	╬		+	\$	Condition	Т	2106c06.51 Soares pregnant uterus NoHPU Homo sapiens cDNA clone
	_						80	491530 3' similar to WP:ZK652.2 CE00448
AATTTTAGOOTEO	H540621	9	3 10	6	28	Examples L21950		Human peripheral benzodiazepine recepior related nixyra
CA166AA1111A1AA		\vdash	L		\vdash		M36035	Human peripheral benzodiazepine receptor (upos) incara
	H540673	-	2 10	~	12	No Match		(CEA B)
CATGGACAAAAAAA	H545152		0	=	7	Examples U19718		Human microfibril-associated glycoprotein (Mr. A. 2).
CATGGACCACCIIIA	H\$45430		0	2	18	Examples M75165		H. sapiens epithelial tropomyosin (1M1) mkwA
CATGGACCAGGCCC		+		-	\vdash		M12125	Human fibroblast muscle-type tropomyosun muscle
		+	T	+	-		M74817	Human tropomyosin-1 (TM-beta) mRNA, complete cas
	050777	-	0	19	2	Examples M74092	M74092	Human cyclin mRNA
CATGGACCCCAAGGC	11646710	1 =	Ľ	=	2	Examples L37033	L37033	Homo sapiens FK-506 binding protein homologue
CATGGACCCTGCCCT	1240/10	+	1	+	+			2b37g02.s1 Soares parathyroid tumor NoHPA Homo sapiens CDINA Clone
	H548062	-	- -	13		Examples N90046	N90046	305810 3'
W CAL SAMOOT AT CLOS		\vdash		-	-			2106a10.s1 Soares pregnant uterus ivoru o monto sapiens com e como
							AA115048	491514 3'
99409090400#4	H551315	~	4 5	32	3	Examples M63193	M63193	Human platelet-denved endothelial cell growni lactor
ののなっていることでは、 かったいましましましましましましましましましましましましましましましましましましまし	H554876	=	3	0	14	Examples M61764	M61764	Human gamma-tubulin mKNA,
していまってつなってい	H559615	0	0	2	01	Examples D17793	D17793	Human mKNA (HA1/33) for OKF
CELECTOR OF CELECT	H\$60056	0	8	32	E	Examples S68252	S68252	TIMP-1=metalloprotemase inhibitor
(A) GGAGAGIGIGIG		+			-		X02598	EPA glycoprotein (erythroid-potentiating activity)
		+	L		\vdash		X03124	tissue inhibitor of metalloproteinase 2
1) CATGGAGCAGGATGA	H561807	0	0	-	12	No Match		
	716/7406			4	-2	Examples	Examples AA214523	2r89c01.s1 Soares NbHTGBC Homo sapiens cDNA clone 682848 3'
1) CATGGAGGGAGTTCC	H30/400	+	_	1.	+		N30324	yw75d01.s1 Homo sapiens cDNA clone 258049 3'
	1870787	10	2	İ	2	Examples X70070	X70070	H.sapiens mRNA for neurotensin receptor.
UNICATGGAGTCCGGAGC	1,0/0/61	5		•				

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2612c08.s1 Soares fetal heart NbHH19W Homo sapiens cDNA clone 358766 3' similar to SW:YA94_SCHPO Q09783 HYPOTHETICAL 11.4	KD PROTEIN C13G6.04 IN CHROMOSOME 1	2k72d06.s1 Soares pregnant uterus NbHPU Homo sapiens cDNA clone		Г	zk46c12.s1 Soares pregnant uterus NbHPU Homo sapiens cDNA clone		methyl sterol oxidase (ERG25)		Human mRNA for elongation factor-1-beta.	H.sapiens mRNA for elongation factor 1-beta	Human nicotinamide N-methyltransferase (NNMT) mRNA, 0	Human mBNA for 14kDa beta-galactoside-binding lectin	Luman mPNA for heta-palactoside-binding lectin	Usuna 14 kd ledin mRNA complete cds	mulian 14 na total inter 4 companies	HL14=0cta-galactostuc ontonic process	489319 5 Similar to Columnia And Teperate Series CDNA clone 668614 3	similar to gb:X02492 INTERFERON-INDUCED PROTEIN 6-16		Human mRNA for pancreatic carcinoma marker GA/33-1, 0		55 (491117.3)	2170h04.s1 Stratagene colon (#937204) Homo sapiens cDNA clone 510007	Т	Tumman usemlar endothelial growth factor B 186	Human cytochrome c oxidase subunit VIb	Human histone H1 (H1F4) gene, complete cds
	W94333		A046631	R91942		AA040439	J60205		K 60489	X60656	10801	215356	0007	V1407)	J04430	244881	AA05448		AA24372	X13425		AA13698		AA05334	043368	M38759	M60748
	No Match	100	Examples AA046631				Examples U60205	No Match	Examples X60489	^	Example: 1108071	2	Examples A13230		7		Examples AA054483		Examples AA243725	Examples X13425		Examples AA136985		Examples AA053346	Examples U43368	D22019	Evaniphe
	900	7	19	-	╁		12	88	25		-	1	7	7	1	1	 16		~	39		0		33		٩	2
	1		7		\dagger			73	_		٤	_1_	7				 ~		2 44	4 23		5 27		- 1	2	1	207
	,	<u>`</u>	5	L	1		0 5	1	10	1		5	4	-			 -	-	_ <u>~</u>	١.	-	∞		29 24	9	- 1-	× -
	1	+	~	+	+		-		_		 	╛	=	-		_	0		00	0		6		12	-	1	
		H572806	H585913	21,000			H587800	H589825	H605956			H606471	H611597				H616224		HK17891	H618841		H633577		H643707	H655177		H655361
		95 CATGGAGTTCGACCT		96 CATGGATTAAGTGAG				97 CATGGAT TGACCIC	98 CATGGCAAAAAAA	99 CATGGCATTIAMIA		100 CATGGCCAACAACGA	101 CATGGCCCCCAATAA							03 CAT GGCCG 1 CGGAGG	מין כאן פפרכן אככרפאפ	105 CATGGCGGGGTGGAG		 CATGGCTCAGCTGGA	117 CATGGCTTTTCAGAC		1118 CATGGGAAAAAAA

		_	_	_	-	-	- //TY//AT	
		+	+		+			
	17665547	×	13	Ŕ	-	Examples X02920		Human mRNA for alpha 1-antitrypsin carboxyterminal, 0
CATGGGAAAAGTGGT	TCCCOU.	_	_		1			Human mRNA for alpha 1-antitrypsin
		+	+				V00496	Human messenger RNA for alpha-1-antitrypsin
		+	+				100001	Human alpha-1 antitrypsin gene, 3' end
		+	+	1	T			zi22b01.s1 Soares pregnant uterus NbHPU Homo sapiens cDNA clone
	H658059	_	- 4	Ø	16	Examples	Examples AA127040	502633 3'
110 CATGGGAAGGGAGGC	COOCOLI	+	L	L			Г	zd86f06.s1 Soares fetal heart NbHH19W Homo sapiens cDNA ctone
							W81387	347555 3'
		+	+	T	T		H45477	
	11000043	+	4	19	12	Examples D26598		Human mRNA for proteasome subunit HsC10-II. , 0
CATGGGAGTCATTGT	H000947	=	L	1	1=	Examples N74310		za78c01.s1 Homo sapiens cDNA clone 298656 31
112 CATGGGAGTGTGCGT	H00/30/	+	 	1	1			yt92e01.s1 Homo sapiens cDNA clone 231768 3'
		-	1				700762	sen 2272 Homo saniens cDNA clone ssb4HB3MA(extended-ft-6) 3'
		+	- 1:	1	1	Examples X17567		H. sapiens RNA for snRNP protein B
CATGGGATTGTCTGG	H671455	1		1_	1			Human small nuclear ribonucleoprotein particle SmB
	000000000000000000000000000000000000000	-	5	0	12	Examples M69054		Human insulin-like growth factor binding protein 6, 0
CATGGGCCCCTCACC	H6//320	7						Human insulin-like growth factor binding protein 6
	1577751	10	- -	7	7	Examples N74323		za78d08.s1 Homo sapiens cDNA clone 298671 3
CATGGGCCCTCTGAG	2	+	+					yo18f08.s1 Homo sapiens cDNA clone 178311 3
		\dagger	+				H41102	yn88a08.s1 Homo sapiens cDNA clone 175478 3'
		\dagger	+					zm84b09.s1 Stratagene ovarian cancer (#937219) Homo sapiens culvA
	H686815	0	-	3 13	22	Examples	Examples AA074777	clone 544601 3'
1:000010000		\dagger	-					zm04a04.s1 Stratagene corneal stroma (#93/222) riolliu sapielis CD140
							AA062735	clone 513102 3'
		\dagger	+	1				zm63f12.s1 Stratagene fibroblast (#937212) Homo sapiens cDNA clone
							AA112905	530351 3'
	H688713	25	-	0	72	No Match		
CALGGGGAAGCACA	H690863	7	_	1 16	7	No Match		
CALGGGGAGGGGGGGGGGGGGGGGGGGGGGGGGGGGGGG	H690890	1-	0	1 14	_	No Match		44 44
119 CATGGGGAGGTAGCA	11693117	=	L	3 39	2	Examples V00523	V00523	Human mRNA for histocompatibility antigen HLA-DR
120 CATGGGGCATCTCTT	1102011	1	╀				X00274	Human gene for HLA-DR alpha heavy chain a class II
		1	\dagger	1			1711171	Himan H. A-DR alpha-chain mRNA

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		\vdash		-	\vdash		100202	human hla-dr heavy chain gene; 3' ilank
E 400 000 000 000 000 000 000 000 000 00	1715401	╞	0	2	7	Examples U18009		Human chromosome 17q21 mRNA clone LF113.
CATGGGTGGGGAGA1	,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,	╀	- I	+	┝			EST57778 Homo sapiens cDNA 3' end similar to None
		+	1	+	H		T33339	EST57474 Homo sapiens cDNA 3' end similar to None
A CONTROL OF THE CONT	H778778	-	3	191	8	Examples M59911		Human integrin alpha-3 chain mRNA
CATGGIACIA	H778810		10	15	S	Examples X87689		H.sapiens mRNA for putative p64 CLCP protein
CALGELACTET GCC1	H737344	1	1	2	-	Examples L12350		Human thrombospondin 2 (THBS2) mRNA
124 CATGGT CAAAA111C		1_	4	79	62	Examples D21261		Human mRNA (HA1756) for ORF
25 CATGGTCTGGGGC11				+	\vdash			Human keratinocyte cDNA, clone 686
	H752521	-	5 7	12	77	Examples H51290		yp07a05.s1 Homo sapiens cDNA clone 186704 3'
26 CATGGTCTGTGAGAG	11102021	+		+	-			yx44g12.s1 Homo sapiens cDNA clone 264646 3'
		+	1	+	ŀ			2076e09.s1 Stratagene pancreas (#937208) Homo sapiens cDNA clone
							AA158271	592840 3'
99 なしつおうおうかんかん ここ	H752531	6	0	-	13	No Match		
CONTROLLES OF THE CONTROL OF THE CON	H753162	0	1 2	=	2	No Match		
(28 CATGGTCT TGARGCC	FCEPSCH	1_	14 42	13	68	Examples X87373		Class C, H.sapiens RPS3a gene
129 CATGGTGAAGCCAGI	1754567 1754567	١	_	┢	흐	Examples X08058		GLUTATHIONE S-TRANSFERASE P (HUMAN)
CATGGT GAATGACGG	H760361	6	3	F	25	Examples X51439		Human mRNA for serum amyloid A (SAA) protein
3 CATGGTGCGGAGGAC	1000011	,	L		96	Examples U15008		Human SnRNP core protein Sm D2 mRNA
132 CATGGTGCTGGAGAA	H/01401	╬		: -	i z	Examples U62800		Cystatin M (CST6)
CATGGTGGAGGCAC	1765003	1 2	. 5	5	É	Examples H46430		yo12h12.s1 Homo sapiens cDNA clone 177767 3'
114 CATGGTGGTACAGGA	COOCO/ L	=	-	1	+			zf13a06.s1 Soares fetal heart NbHH19W Homo sapiens cDNA clone
							AA047563	376786 3'
		╁	+		\vdash			zo13f02.s1 Stratagene colon (#937204) Homo sapiens cDNA clone 380/19
							5	3,
でなりの事がなりませいの事なから、	H774629	0	7	13	6	Examples X59288		H. sapiens gene for intercellular adhesion molecule
CALGGI ICACIGCAG		\dagger	-					Human major group rhinovirus receptor (HRV) mRNA
		+	-		\vdash		103132	Human intercellular adhesion molecule-1 (ICAM-1)
		+	╀		-		M55100	Human cell surface glycoprotein P3.58 mRNA
	H781823	┢	1	õ	24	Examples K02765	K02765	Human complement component C3 mRNA, alpha and beta
A STATE OF THE CONTROL OF THE CONTRO		178 110		14 340	139	Examples M17987	MI7987	Human beta-2-microglobulin gene
138 CATGGTTTAAATCGA	H782391	-	6 12	4	14	Examples D00760	D00760	Human mRNA for proteasome subunit f10.3
	07170711	-	9		12	Examples X57025	X57025	INSULIN-LIKE GROWTH FACTOR IA PRECURSOR (HUMAN)
139 CATGTAGGCT LAAC	U807703	10	1_	7	þ	No Match		
1.10 CATGTAATTTTGGAA	110021	,]				

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E & COMMUNICATION OF THE PARTY	H802793		\vdash	No Match	ch	
CATGTAATTTTGGAL	110020011	7	٣	14 Examp	Examples X85373	H. sapiens mRNA for Sm protein G
11 CATGTACATTTTCAT	10003370		0	_	ch	
CATGTACCCCGTACA	0 2/00/01/	10	12		ਚ	
1) CATGTACCCTTCTAT	100001	L	L	74 Examp	Examples 102931	Human placental tissue factor (two forms) mRNA
11 CATGTAGGAAAGTAA	H87/43/			L	M16553	Human tissue factor mRNA, complete cds
		T	+		M27436	Human tissue factor gene, complete cds
	21715011	17	08	130 Examp	Examples X64899	H. sapiens mRNA homologous to mouse P21 mRNA.
15 CATGTAGGTTGTCTA	H851410 42				X16064	Human mRNA for translationally controlled tumor protein
			-			riedway account bellanders
-					L13806	Homo sapiens (clone 04) translationally controlled tumor protein
	1 1830677	0	- - -	16 Examp	Examples M98479	Human transglutaminase mRNA
	H851834 0	L	191	3 Examp	Examples D12149	Human HepG2 3'-directed Mbol cDNA, clone 5247
CATGTALTICISCO	01 00035011	76 26	24	48 Examp	Examples X80909	H.sapiens alpha NAC mRNA
1N CATGTCACAAGCAAA	07 07507011		43	_	Examples X56134	Human mRNA for vimentin.
19 CATGTCCAAATCGAT	Н808ЭОУ	<u>}</u>	7		719554	H.sapiens vimentin gene
		1	$\frac{1}{1}$		M14144	Human vimentin gene, complete cds
		+	1		M25246	Human vimentin (HuVim3) mRNA, 3' end
	U870310	0	12	2 Exam	Examples N92906	zb57a08.s1 Homo sapiens cDNA clone 307670 3'
AN TATGTCCACTGGCCT	1	<u>` </u> 				
					T17488	NIB978 Normalized infant brain, Bento Soares Homo sapiens cDNA 3'end
		\vdash			AA349906	EST56900 Infant brain Homo sapiens cDNA 3' end
	H871920 6	6 10	25	5 Exam	Examples X67016	H. sapiens mRNA for amphiglycan
CATGLCCATCIGITG	1	1_	<u>. </u>		D13292	Human mRNA for ryudocan core protein
	H899060	5 15	-	69 Exam	Examples M77233	Human ribosomal protein S7 mRNA
CATGLCGICTITATE	L	5 2	46	19 Exam	Examples S48568	tissue inhibitor of metalloproteinase 2 (3'-end region)
ייי האופורורופאופרי		-				
OTO A TOTTOTOTAN	H916232 0	4 3	-		Examples N71680	yz93b03.s1 Homo sapiens cDNA clone 2905/3 3
155 CATGTCTTGTGCATA		22 15	20	45 Exam	Examples X03083	Human lactate dehydrogenase-A gene
		_			X02152	Human mkny ior iacuse uchyu ogchasory
	1				X02153	Human pseudogene for lactate denydrogenase-A
136 CATICACTG	H920392	1	0 9	16 No Match	atch	
	_		7	11 Exam	Examples X07979	CTGTGG, Class A, Human mRNA for fibronectin receptor beta subunit.
157 CATGTGAAGTTATAC	n 1c7c076H			1		

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		-		+	-	-			zkośho7.s1 Soares pregnant uterus Nbł.PU Homo sapiens cDNA clone
		H012731	- 80		=	12 E	Examples AA027860		469693 3'
ISS CATGTGATGTCTGG	+	H018876	2 -			١	Examples M25753		G2/MITOTIC-SPECIFIC CYCLIN BI (HUMAN)
150 CATGTGCCATCTGTA	+			-					yc22c04.s1 Homo sapiens cDNA clone 81414.3
		+		\perp	-		14	R67969	yi29g08.s1 Homo sapiens cDNA clone 140/02 3
					_				2.01 ft 3 s 1 Stratagene ovarian cancer (#937219) Homo sapiens cDNA
									Clone \$94269 3' similar to SW: NGAL HUMAN P80188 NEUTROPHIL
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i 160 CATGTGCCCTCAAAA		H939841 1	=	7	2	2 2	- Yaminay	1	2h15d08 s1 Homo sapiens cDNA clone 302127 3' similar to
									SW.NGAL HUMAN P80188 NEUTROPHIL GELATINASE-
		0700001	7	_		19	Examples N79823		ASSOCIATED LIPOCALIN PRECURSOR
161 CATGTGCCCTCAGAA		242		1					
									zm90h04.s1 Stratagene ovarian cancer (#937219) Homo sapiens cDNA
									clone 545239 3. similar to 5 W. N. CALL, ILONARY CONTROLL OF A TIMA SP. A SSOCIATED LIPOCALIN PRECURSOR
162 CATGTGCCCTCAGGA	-	H939851 1	13 31	9	22	83	xamples	Examples AAU / 3890	מניתו וועיתו המספים ביים
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		H941856	3	1	2	12 E	3xamples	Examples AA100279	3.
(a) CA161 GCC1 131CC	<u> </u>	H044038	2 5	2	17	<u>~</u>	No Match		Nation Conject Clone
104 CATGTGCGCTGGCCC	+		L	L	-	-			zk10a01.s1 Soares pregnant uterus north o nomo saproms contra como
<u> </u>		0230701	- 4	ν.	4	16 I	Examples	Examples AA029262	470088 3'
165 CATGTGCTTCATCTG	1	000	1	1	L				yv66e10.s1 Soares fetal liver spleen INFLS Homo sapiens cDNA clone
								N54281	247722 3'
		+	+	†	+	+			zn76c02.s1 Stratagene NT2 neuronal precursor 937230 Homo sapiens
								AA114075	cDNA clone 564098 3'
	+	13053051	18	1	22	48	Examples L76200	L76200	Homo sapiens guanylate kinase (GUK1) mRNA
Ind CATGTGGAGTGGAGG		1070	- [-	1 2		Examples X00570	X00570	Human mRNA for precursor of apolipoprotein CI
167 CATGTGGCCCCAGGT	1	7000701	1 2	1-	1/2	L	Examples L16510	L16510	Homo sapiens cathepsin B mRNA
168 CATGTGGGTGAGCCA	1	H90706H	.1_		1			M14221	Human cathepsin B proteinase mRNA, complete cds
		1076446	1"	٣	15	~	Examples L35240	L35240	Human enigma gene
169 CATGTGTGAGCCCCT		266440	1	1	 ≃		Examples L38941	L38941	Homo sapiens ribosomal protein L34 (RPL34) mRNA
170 CATGTGTGCTAAATG	-	H9/0044	1		2,4		Fxamples	X03473	Human gene for histone H1(0).
1-1 CATGTGTGTGTTTGT	-	H978687	-		+				zk23g08.s1 Soares pregnant uterus NbHPU Homo sapiens cDNA clone
		H007044			21	-	Examples	Examples AA034505	471422 3'
1-2 CATGTTATGGATCTC	-	4	,	1		1			

zt31b06.s1 Soares ovary tumor NbHOT Homo sapiens cDNA clone 723923	3'	ZK30c10.s1 Soares pregnant uterus NoHPU Homo sapiens CDINA Cione	20404 of Home conjune cDNA clone 236071 3'	yusouuv,si muino sapiens enitti come urran X12	EST04595 Homo sapiens culva cione in Drast	Source Homo saniens cDNA	NIBIONA NOTHIGHT COLUMN	3'end similar to ESTU4575 m. Saprens CONTROLL CONTROLL	2e97h02.s1 Soares fetal heart NbHH19W Homo sapiens CDINA CIONE	366963 3'	Sansas NhHTGBC Homo saniens cDNA clone 712204 3	CIUDAUDISI DOMESTICA ALMA ALMA AKK75 31	ym05a09.s1 Homo sapiens civing Hours 400.5.3	H. sapiens mRNA for tyrosine kinase receptor.	Human mRNA for collagen VI alpha-1	H sapiens gene for glutaminyl-tRNA synthetase	7473110 s1 Soares pregnant uterus NoHPU Homo sapiens cDNA clone	488415 31	vz36b07.s1 Homo sapiens cDNA clone 285109 3'		zt71g03.s1 Soares testis NHT Homo sapiens cDNA clone 727828 3'	H.sapiens (5) Ferritin H pseudogene.	Human mRNA for apoferritin H chain type	Human apoferritin H gene exons 2-4	Human ferritin heavy chain mRNA, complete cds	Human ferritin heavy chain mRNA, complete cds	Human interferon-inducible mRNA (cDNA 6-26).	Human promyelocytic leukemia cell mRNA	Human thymosin beta-4 mRNA, complete cds	zb17a08.s1 Homo sapiens cDNA clone 302294 3'	2133d02.s1 Soares ovary tumor NbHOT Homo sapiens cDNA clone 724131			347396 3'
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				H1003443						H1014660				720100111	H1021270	H1023520			H1024568			7.0000	7 +180701H				303750111	H102/323			H1037777	``		
				ASPERTMENT AGA	ישופון וכעון ופועי						1 CATGTTC1616AA1C				S CATGCCCCCTG	1-10 ATGTTGCTGACTTT			TT CATGTTGGAGATCTC				"N CATGTTGGGGTTTCC					179 CATGTTGGTGAAGGA			IND CATGITICCCICAAA			

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		LICENTIFICATION OF THE COLUMN TO THE COLUMN	10101			IN TOPAGETT GCACCT IT				- COLONDOLL COLO					

Table 5 - Transcripts increased in pancreas and colorectal cancer

SAGE tag that were elevated in both in coloreactal and pancreatic tumor,

and are likely to be specific for tumor in general.

Tog Canience		Tag	Tag Number Accession	Accession	HOH
1 ag_ Sequence	טמט		-950498 M10629	10629	d with polyA
CAIG	1	-	-294155 042376	42376	Human retinoic acid induced RIG-E precursor (E) mK
2 CATG CACIICAAGG		+	2	056145	Human thymic shared antigen-1/stem cell antigen-2
San annual or and a	١	(A)	-243747 303040	03040	Human SPARC/osteonectin mRNA, complete cds.
SCAIG AIGIGA	1	_	Σ	M25746	1.
CASSAGOOD STACK	CAC	-	-610466 X53416	53416	Human mRNA for actin-binding protein (filamin) (AB
	TAC		-229106 X02761	19720	ا ـ
מוכיוום אוכיוום	200	-	×	K00799	fn) 3' coding region a
SASTOSOSOTS STANKS	SAG C		-760291 X58536	58536	Human mRNA for HLA class I locus C heavy chain.
			2	M26432	gene, comprere cas:
PATOPROGETTACE	ACG G	_	-76231 M95787	195787	Human 22kDa smooth muscle protein (SMZZ) mknA, com
2000		-		M83106	Human SM22 mRNA, 5' end.
TOTAL CTATO	TGT A	-	-769020 M77349	177349	
	C 860		-589267 X53279	(53279	Human mRNA for placental-like alkaline phospharase
אַראופ פאוווכן	202	-		X55958	atase.
		-		304948	
	£ 0,0±0	-	-85882 X57351	X57351	induci
10 CATG ACCATICTES	, ,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,	1		X02490	Human interferon-inducible mRNA (cDNA 1-8).
	200	-	-884181X15804	X15804	Human mRNA for alpha-actinin.
TICATE ICCITCION	E 0	E	-515821 D80012	080012	Human mRNA for KIAA0190 protein.
IZICATE CITCIGIGIA		+	-241665 M74090	M74090	
13 CATG ATGTAAAAA		+		303801	Human lysozyme mRNA, complete cds with an Alu repe
		+		M19045	e cds.
ORDBADADED DEAD FI	CONCE	+	-673954 X17620	X17620	Human mRNA for Nm23 protein, involved in developme
				X75598	
ASASTE ANTATIGAGA	GAGA	-	-53129 062962	062962	
16 CATE TTTTGATAA		-	-1048113 016891	016891	region cDNA,
ACCEPTORATION OF ACTION	1	-	-302741 X53743	X53743	H. sapiens mRNA for fibulin-1 C.
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CATG ACATCATCGA T		00000	usms earlens (clone 01) liver expressed protein mR
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CATG CTGTTGGTGA T507577 1514530 Human mRNA for ribosom 2	ACATCATCGA	L06505	protein S28,
CATG ATTATTTC T	CATG CTGTTGGTGA	П	ein L7.
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CATG CTCTTCCAGA A	CATG GGCDGGGG	L25346	sapiens ribosomal protein L2/ (nomologue of
Y00483 Human gene for gluthathlook	CATE CTCTTCGAGA	-490889 Y00433	1 1 1 1
X13710 H.sapiens unspliced mRNA X13709 Human gpx1 mRNA for gluth X13709 Human gpx1 mRNA for gluth M21304 Human glutathione peroxid M21304 Human glutathione peroxid U00947 Human liver mRNA fragment U00947 Human clone C4E 3.2 (CAC) U00947 Human clone C4E 3.2 (CAC) U00947 Human clone C4E 3.2 (CAC) U00947 Human mRNA for LLRep3. L21756 Human mRNA for Epstein-Ba L21756 Human mRNA for HBpl5/L22, U1752 Human mRNA for HBpl5/L22, U1752 Human mRNA for HBpl5/L22, U1752 Human mRNA for HBpl5/L22, U1753 Human mRNA for elongation U23765 Human mRNA for elongation U23765 Human mRNA for elongation U1785 Human HepG2 3' region U1785 U1785 Human HepG2 3' region U1785		Y00483	2
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		017276	3' region Mbol cDNA, clone

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		mKNA, complete
	L41498	Complete
A CATACCATATC A	-988366 057846	Human ribosomal protein L39 mRNA, complete cus.
DESCRIPTION OF ACTION OF A	-621035 X71973	H. sapiens GPx-4 mRNA for phospholipia nyaroperoxic
1	-383489 226876	H. sapiens gene for ribosomal protein L30.
	-803369 X69391	H. sapiens mRNA for ribosomal protein Lb.
Thermore	-803369 D17554	AKEDIO',
	-803369 S71022	neoplasm-related C140 product [human, tnyroid care
SO CATE AACGACCTCG T	-24951 V00598	Human beta-tubulin pseudogene.
	-24951 V00599	prote
SACATG CCCTGCCTTG T	-358783 X55110	mRNA for neurite outgrowth - promotting
CATG	-346761 038846	of the binding (5:2)
	D16933	Human HepG2 3' region count, crouc increase
SS CATG AGCACCTCCA G	-148949 211692	וסוואמרדסוו דמככס
SECATG CGCCGGAACA C	-416261 X73974	H. sapiens HREL4 manna.
	053660	mRNA for ribosomar process,
57 CATG CTAAAAAAA A	-458753 M33680	26-KDa Cell Sullace Process
CATG	-686319 009510	Syllinecase mens complete
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	D30658	Tototo bomologi
59 CATG ATTCTCCAGT A	-253260 X55954	.
	x52839	Human mRNA for ribosomar process brokein.
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	X15005	Human mkNA Ior potential raminist consor
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	103799	י <u>ה</u>
	M14199	Complete
61 CATG CAGCTCACTG A	-302367 D87735	anadmon 'E
	L10376	Human (Clone Cid-Bos) many sequences:
	880520	CIECUTAE TEPERE CONTRACTOR
G CHARATTER G	-200576 014973	

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			1	131610	Home sapiens (clone cori-1c15) \$29 ribosomal prote
			1	Т	and one man for ribosomal protein 18.
63	63 CATG AATCCTGTGG	A	-5522/22840/	7	H. Saptells mines for month complete cds.
6.4	CATG AATAGGTCCA	A	-51925 M64716	╗	
,		() F	-1-	-1 X83412	H.sapiens Bl mRNA for mucin.
65	CATG AAAAAAAAA	~ 1	2	Г	pp) for
			2		receptor (81/
			×	X76180	H.sapiens mRNA for lung amiloride sensitive Na+ ch
				008470	Human FR-gamma' mRNA, complete cds.
				U08471	or 3 mRNA, complete cds.
				048697	
				D28532	phospha
			-	M55914	Human c-myc binding protein (MBP-1) mRNA, complete
				1.06175	
		-		573775	_,
				877393	transcript ch138 [human, RF1, RF48 stomach cancer c
į				x60036	H sabiens mRNA for mitochondrial phosphate carrier
		,	235645 X79238	779738	H sapiens mRNA for ribosomal protein L30.
99	CATG CCAGAACAGA		108666	116991	Human thymidylate kinase (CDC8) mRNA, complete cds
			200000	20000	H sapiens mRNA for ORF.
67	67 CATG AAGGTGGAGG	A	-44083	770004	
68	68 CATG CCTAGCTGGA	£.	-379369 X52856	0087CX	processed
				X5285/	Cyclophizzzi zered recessed
				X52854	Human cyclophiliniterated processes From 5.2.1.8
				X52851	TOT CYCLOPHICE CO.
				Y00052	
18	SO CATG GAACACATCC	A	-528694 X63527	X63527	
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	M26325	Human cytokeratin 18 mRNA, 3' end.
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	-177315 D86979	Human male bone marrow myeloblast mknA for himhore
	X55923	Human DNA for Alu element PIN6.
	66967X	H. sapiens ALU repeat, 230bp.
	X12544	Human mRNA for HLA class II DR-Deta (HLA-DK B).
	277989	Hindili II
	011831	Human clone 2102V-I chromosome 18p telomeric seque
	1112580	Human Alu repeat sequence A3.
	012582	Human Alu repeat sequence B2.
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	1114696	
	111 4 697	Human Alu-Sb2 repeat, clone HUM-11.
	1114698	Human Alu-Sb2 repeat, clone HSB-8P.
	111 4699	Alu-Sb2
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	014701	Human Alu-Sb2 repeat, clone HSB-2P.
	014704	Alu-Sb2 repeat,
	014706	Alu-Sb2 repeat,
	014707	clone HUM-7.
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Isolation of partial cDNA (3' fragment) by 3' directed PCR reaction

This procedure is a modification of the protocol described in Polyak et al. (1997) Nature 389:300. Briefly, the procedure uses SAGE tags in PCR reaction such that the resultant PCR product contains the SAGE tag of interest as well as additional cDNA, the length of which is defined by the position of the tag with respect to the 3' end of the cDNA. The cDNA product derived from such a transcript driven PCR reaction can be used for many applications.

RNA from a source believed to express the cDNA corresponding to a given tag is first converted to double-stranded cDNA using any standard cDNA protocol. Similar conditions used to generate cDNA for SAGE library construction can be employed except that a modified oligo-dT primer is used to dreive the first strand synthesis. For example, the oligonucleotide of compositon 5'-B-TCC GGC GCG CCG TTT T CC CAG TCA CGA(30)-3', contains a poly-T stretch at the 3' end for hybridization and priming from poly-A tails, an M13 priming site for use in subsequent PCR steps, a 5' Biotin label (B) for capture to strepavidin-coated magnetic beads, and an AscI restriction endonuclease site for releasing the cDNA from the streptavidin-coated magnetic beads. Theoretically, any sufficiently-sized DNA region capable of hybridizing to a PCR primer can be used as well as any other 8 base pair recognizing endonuclease.

cDNA constructed utilizing this or similar modified oligo-dT primer is then processed exactly as described in U.S. Patent No. (insert) up until adapter ligation where only one adapter is ligated to the cDNA pool. After adapter ligation, the cDNA is released from the streptavidin-coated magnetic beads and is then used as a template for cDNA amplification.

Various PCR protocols can be employed using PCR priming sites within the 3' modified oligo-dT primer and the SAGE tag. The SAGE tag-derived PCR primer employed can be of varying length dictated by 5' extension of the tag into the adaptor sequence. cDNA products are now available for a variety of applications.

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This technique can be further modified by: (1) altering the length and/or content of the modified oligo-dT primer; (2) ligating adaptors other than that previously employed within the SAGE protocol; (3) performing PCR from template retained on the streptavidin-coated magnetic beads; and (4) priming first strand cDNA synthesis with non-oligo-dT based primers.

Isolation of cDNA using GeneTrapper or modified GeneTrapper Technology

The reagents and manufacturer's instructions for this technology are commercially available from Life Technologies, Inc., Gaithersburg, Maryland. Briefly, a complex population of single-stranded phagemid DNA containing directional cDNA inserts is enriched for the target sequence by hybridization in solution to a biotinylated oligonucleotide probe complementary to the target sequence. The hybrids are captured on streptavidin-coated paramagnetic beads. A magnet retrieves the paramagnetic beads from the solution, leaving nonhybridized single-stranded DNAs behind. Subsequently, the captured single-stranded DNA target is released from the biotinylated oligonucleotide. After release, the cDNA clone is further enriched by using a nonbiotinylated target oligonucleotide to specifically prime conversion of the single-stranded target to double-stranded DNA. Following transformation and plating, typically 20% to 100% of the colonies represent the cDNA clone of interest. To identify the desired cDNA clone, the colonies may be screened by colony hybridization using the 32P-labeled oligonucleotide as described above for solution hybridization, or alternatively by DNA sequencing and alignment of all sequences obtained from numerous clones to determine a consensus sequence.

The genes which are identified herein as being differentially expressed in normal and cancer cells can be used diagnostically and prognostically. Transcription levels in a test sample suspected of being neoplastic can be determined and compared to the levels in normal colon cells. The test sample may be from any tissue suspected of neoplasia, and particularly from either suspected colorectal or suspected pancreatic cancer cells. The control cells for

the purposes of comparison are normal cells, preferably of the same tissue type as the test sample, e.g., colon cells, or pancreatic duct epithelial cells. Upregulation of transcription or downregulation of transcription is therefore diagnostic of the neoplastic state, depending on what gene is used as a test reagent. Similarly, transcription levels can be monitored to assess patent responses to anti-tumor therapies. Transcription levels will also provide prognostic information. For example, the level of transcription in a test sample can be compared to levels found in bona fide normal and tumor cells. More extreme deviations from normal expression levels indicate a poorer prognosis.

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Transcription levels can be determined according to any means known in the art. These include, without limitation, Northern blots, nuclear run-on assays, in vitro transcription assays, primer extension assays, quantitative reverse transcriptase-polymerase chain reactions (RT-PCR), and hybrid filter binding assays. These techniques are well known in the art. See J.C. Alwine, D.J. Kemp, G.R. Stark, *Proc. Natl. Acad. Sci. U.S.A.* 74, 5350 (1977); K. Zinn, D. Di-Maio, T. Maniatis, *Cell* 34, 865 (1983); G. Veres, R.A. Gibbbs, S.E. Scherer, C.T. Caskey, *Science* 237, 415 (1987).

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Similarly, upregulated genes and downregulated genes can be detected by measuring expression of their protein products. This can be done by any means known in the art, including but not limited to Western (immuno) blot, enzyme linked immunoadsorbent assay, radioimmunoassay, and enzyme assay. Such techniques are well known in the art. Protein products can be detected in tissue samples of a test patient, using a suspect sample as a test sample, and a matched normal tissue sample from the same tissue type as a control. If normal tissue is not available then a closely related tissue type can be used. Desirably both the samples being compared will be from the same individual. Alternatively, aberrant expression levels of protein products can be detected in body samples, such as blood, serum, feces, urine, sputum. As a control, a normal matched sample can be used from a healthy individual. Aberrant expression levels of transcripts can also be detected in such body samples, particularly in blood and serum.

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Probes for use in the assays for transcription levels of particular genes or sets of genes may be RNA or DNA. The probes will be isolated substantially free of other cellular RNAs or DNAs. If the reagent contains one probe then it will comprise at least 50% of the nucleic acids in the reagent composition. If the reagent contains more than one probe, then the proportion will decrease accordingly, so that specific probes will still comprise at least 50% of the nucleic acids in the reagent composition.

Probes can be labeled according to any means known in the art. These may include radioactive labels, fluorescent labels, enzymatic labels, and binding partner labels such as biotin. Means for labeling and detecting probes are well known in the art. Probes comprise at least 10, 11, 12, 15, 20, or 30 contiguous nucleotides of a selected gene.

This invention provides proteins or polypeptides expressed from the polynucleotides of this invention, which is intended to include wild-type and recombinantly produced polypeptides and proteins from procaryotic and eucaryotic host cells, as well as muteins, analogs and fragments thereof. In some embodiments, the term also includes antibodies and anti-idiotypic antibodies.

It is understood that functional equivalents or variants of the wild-type polypeptide or protein also are within the scope of this invention, for example, those having conservative amino acid substitutions. Other analogs include fusion proteins comprising a protein or polypeptide.

The proteins and polypeptides of this invention are obtainable by a number of processes well known to those of skill in the art, which include purification, chemical synthesis and recombinant methods. Full length proteins can be purified from a colon or pancreatic cell or tissue lysate by methods such as immunoprecipitation with antibody, and standard techniques such as gel filtration, ion-exchange, reversed-phase, and affinity chromatography using a fusion protein as shown herein. For such methodology, see for example Deutscher et al. (1999) Guide To Protein Purification: Methods In Enzymology (Vol. 182, Academic Press). Accordingly, this invention also

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provides the processes for obtaining these proteins and polypeptides as well as the products obtainable and obtained by these processes.

The proteins and polypeptides also can be obtained by chemical synthesis using a commercially available automated peptide synthesizer such as those manufactured by Perkin Elmer/Applied Biosystems, Inc., Model 430A or 431A, Foster City. The synthesized protein or polypeptide can be precipitated and further purified, for example by high performance liquid chromatography (HPLC). Accordingly, this invention also provides a process for chemically synthesizing the proteins of this invention by providing the sequence of the protein and reagents, such as amino acids and enzymes and linking together the amino acids in the proper orientation and linear sequence.

Alternatively, the proteins and polypeptides can be obtained by well-known recombinant methods as described, for example, in Sambrook et al., (1989), supra, using the host cell and vector systems described above.

Also provided by this application are the polypeptides and proteins described herein conjugated to a detectable agent for use in the diagnostic methods. For example, detectably labeled proteins and polypeptides can be bound to a column and used for the detection and purification of antibodies. They also are useful as immunogens for the production of antibodies as described below. The proteins and fragments of this invention are useful in an in vitro assay system to screen for agents or drugs, which modulate cellular processes.

The proteins of this invention also can be combined with various liquid phase carriers, such as sterile or aqueous solutions, pharmaceutically acceptable carriers, suspensions and emulsions. Examples of non-aqueous solvents include propyl ethylene glycol, polyethylene glycol and vegetable oils. When used to prepare antibodies, the carriers also can include an adjuvant that is useful to non-specifically augment a specific immune response. A skilled artisan can easily determine whether an adjuvant is required and select one. However, for the purpose of illustration only, suitable adjuvants include, but

are not limited to Freund's Complete and Incomplete, mineral salts and polynucleotides.

This invention also provides a pharmaceutical composition comprising any of a protein, analog, mutein, polypeptide fragment, antibody, antibody fragment or anti-idiotipic antibody of this invention, alone or in combination with each other or other agents, and an acceptable carrier. These compositions are useful for various diagnostic and therapeutic methods.

Antibodies can be generated using the proteins encoded by the transcripts identified by the tags disclosed herein. Use of all or portions of the protein as immunogens is routine in the art. Similarly, fusion proteins can be used as immunogens. Antibodies can be affinity purified using the proteins or portions thereof used as immunogens. Similarly, monoclonal antibodies specifically immunoreactive with the protein sequences of the invention can be generated according to techniques which are well known in the art.

Antibodies can be used analytically to quantitate the expression of particular transcripts identified herein as upregulated or downregulated in cancer. In addition, antibodies can be conjugated or non-covalently linked to cytotoxic agents, such as cytotoxins, radionuclides, chemotherapeutic drugs, etc. Such antibodies can be used therapeutically to specifically target cancer cells in which the protein antigens are upregulated. These include the proteins encoded by the transcripts identified by the tags shown in Tables 2, 4, and 5. Means of making such linked cytotoxic antibodies and of administering the same are well known in the art.

Also provided by this invention is an antibody capable of specifically forming a complex with the proteins or polypeptides as described above. The term "antibody" includes polyclonal antibodies and monoclonal antibodies. The antibodies include, but are not limited to mouse, rat, and rabbit or human antibodies.

Laboratory methods for producing polyclonal antibodies and monoclonal antibodies, as well as deducing their corresponding nucleic acid sequences, are known in the art, see Harlow and Lane (1988) supra and

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Sambrook et al. (1989) supra. The monoclonal antibodies of this invention can be biologically produced by introducing protein or a fragment thereof into an animal, e.g., a mouse or a rabbit. The antibody producing cells in the animal are isolated and fused with myeloma cells or heteromyeloma cells to produce hybrid cells or hybridomas. Accordingly, the hybridoma cells producing the monoclonal antibodies of this invention also are provided.

Thus, using the protein or fragment thereof, and well known methods, one of skill in the art can produce and screen the hybridoma cells and antibodies of this invention for antibodies having the ability to bind the proteins or polypeptides.

If a monoclonal antibody being tested binds with the protein or polypeptide, then the antibody being tested and the antibodies provided by the hybridomas of this invention are equivalent. It also is possible to determine without undue experimentation, whether an antibody has the same specificity as the monoclonal antibody of this invention by determining whether the antibody being tested prevents a monoclonal antibody of this invention from binding the protein or polypeptide with which the monoclonal antibody is normally reactive. If the antibody being tested competes with the monoclonal antibody of the invention as shown by a decrease in binding by the monoclonal antibody of this invention, then it is likely that the two antibodies bind to the same or a closely related epitope. Alternatively, one can pre-incubate the monoclonal antibody of this invention with a protein with which it is normally reactive, and determine if the monoclonal antibody being tested is inhibited in its ability to bind the antigen. If the monoclonal antibody being tested is inhibited then, in all likelihood, it has the same, or a closely related, epitopic specificity as the monoclonal antibody of this invention.

The term "antibody" also is intended to include antibodies of all isotypes. Particular isotypes of a monoclonal antibody can be prepared either directly by selecting from the initial fusion, or prepared secondarily, from a parental hybridoma secreting a monoclonal antibody of different isotype by using the sib selection technique to isolate class switch variants using the

procedure described in Steplewski et al. (1985) Proc. Natl. Acad. Sci. 82:8653 or Spira et al. (1984) J. Immunol. Methods 74:307.

This invention also provides biological active fragments of the polyclonal and monoclonal antibodies described above. These "antibody fragments" retain some ability to selectively bind with its antigen or immunogen. Such antibody fragments can include, but are not limited to:

(1) Fab,

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- (2) Fab',
- (3) F(ab')2,
- (4) Fv, and
- (5) SCA

A specific example of "a biologically active antibody fragment" is a CDR region of the antibody. Methods of making these fragments are known in the art, see for example, Harlow and Lane, (1988) supra.

The antibodies of this invention also can be modified to create chimeric antibodies and humanized antibodies (Oi, et al. (1986) BioTechniques 4(3):214). Chimeric antibodies are those in which the various domains of the antibodies' heavy and light chains are coded for by DNA from more than one species.

The isolation of other hybridomas secreting monoclonal antibodies with the specificity of the monoclonal antibodies of the invention can also be accomplished by one of ordinary skill in the art by producing anti-idiotypic antibodies (Herlyn, et al. (1986) Science 232:100). An anti-idiotypic antibody is an antibody which recognizes unique determinants present on the monoclonal antibody produced by the hybridoma of interest.

Idiotypic identity between monoclonal antibodies of two hybridomas demonstrates that the two monoclonal antibodies are the same with respect to their recognition of the same epitopic determinant. Thus, by using antibodies to the epitopic determinants on a monoclonal antibody it is possible to identify other hybridomas expressing monoclonal antibodies of the same epitopic specificity.

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It is also possible to use the anti-idiotype technology to produce monoclonal antibodies which mimic an epitope. For example, an anti-idiotypic monoclonal antibody made to a first monoclonal antibody will have a binding domain in the hypervariable region which is the mirror image of the epitope bound by the first monoclonal antibody. Thus, in this instance, the anti-idiotypic monoclonal antibody could be used for immunization for production of these antibodies.

As used in this invention, the term "epitope" is meant to include any determinant having specific affinity for the monoclonal antibodies of the invention. Epitopic determinants usually consist of chemically active surface groupings of molecules such as amino acids or sugar side chains and usually have specific three dimensional structural characteristics, as well as specific charge characteristics.

The antibodies of this invention can be linked to a detectable agent or label. There are many different labels and methods of labeling known to those of ordinary skill in the art.

The antibody-label complex is useful to detect the protein or fragments in a sample, using standard immunochemical techniques such as immunohistochemistry as described by Harlow and Lane (1988) supra. Competitive and non-competitive immunoassays in either a direct or indirect format are examples of such assays, e.g., enzyme linked immunoassay (ELISA) radioimmunoassay (RIA) and the sandwich (immunometric) assay. Those of skill in the art will know, or can readily discern, other immunoassay formats without undue experimentation.

The coupling of antibodies to low molecular weight haptens can increase the sensitivity of the assay. The haptens can then be specifically detected by means of a second reaction. For example, it is common to use haptens such as biotin, which reacts avidin, or dinitropherryl, pyridoxal, and fluorescein, which can react with specific anti-hapten antibodies. See Harlow and Lane (1988) supra.

The monoclonal antibodies of the invention also can be bound to many different carriers. Thus, this invention also provides compositions containing the antibodies and another substance, active or inert. Examples of well-known carriers include glass, polystyrene, polypropylene, polyethylene, dextran, nylon, amylases, natural and modified celluloses, polyacrylamides, agaroses and magnetite. The nature of the carrier can be either soluble or insoluble for purposes of the invention. Those skilled in the art will know of other suitable carriers for binding monoclonal antibodies, or will be able to ascertain such, using routine experimentation.

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Compositions containing the antibodies, fragments thereof or cell lines which produce the antibodies, are encompassed by this invention. When these compositions are to be used pharmaceutically, they are combined with a pharmaceutically acceptable carrier.

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The present invention also provides a screen for various agents which modulate the expression of a gene in a pancreatic or colon cell. To practice the method in vitro, suitable cell cultures or tissue cultures are first provided. The cell can be a cultured cell or a genetically modified cell in which a trancript from SEQ ID NOS:1-732, or their complements, is expressed. Alternatively, the cells can be from a tissue biopsy. The cells are cultured under conditions (temperature, growth or culture medium and gas (CO₂)) and for an appropriate amount of time to attain exponential proliferation without density dependent constraints. It also is desirable to maintain an additional separate cell culture; one which does not receive the agent being tested as a control.

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As is apparent to one of skill in the art, suitable cells may be cultured in microtiter plates and several agents may be assayed at the same time by noting genotypic changes, phenotypic changes or cell death.

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When the agent is a composition other than a DNA or RNA, the agent may be directly added to the cell culture or added to culture medium for addition. As is apparent to those skilled in the art, an "effective" amount must be added which can be empirically determined. When the agent is a polynucleotide, it may be directly added by use of a gene gun or

electroporation. Alternatively, it may be inserted into the cell using a gene delivery vehicle or vector as described above.

An agent is a potential therapeutic if it alters the expression of gene in the cell. Altered expression can be detected by assaying for altered mRNA expression or protein expression using the probes, primers and antibodies as described herein.

For the purposes of this invention, an "agent" is intended to include, but not be limited to a biological or chemical compound such as a simple or complex organic or inorganic molecule, a peptide, a protein (e.g. antibody) or a polynucleotide (e.g. anti-sense). A vast array of compounds can be synthesized, for example polymers, such as polypeptides and polynucleotides, and synthetic organic compounds based on various core structures, and these are also included in the term "agent". In addition, various natural sources can provide compounds for screening, such as plant or animal extracts, and the like. It should be understood, although not always explicitly stated that the agent is used alone or in combination with another agent, having the same or different biological activity as the agents identified by the inventive screen. The agents and methods also are intended to be combined with other therapies.

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The above disclosure generally describes the present invention. A more complete understanding can be obtained by reference to the following specific examples which are provided herein for purposes of illustration only, and are not intended to limit the scope of the invention.

EXAMPLE 1

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This example demonstrates the characterization of the general transcription of human colorectal epithelium, colorectal cancers, and pancreatic cancers.

We used the recently developed SAGE (serial analysis of gene expression) method to identify and quantify a total of 303,706 transcripts derived from human colorectal (CR) epithelium, CR cancers or pancreatic cancers (Table 1A) (3). These transcripts represented approximately 48,741

different genes (4) that ranged in average expression from 1 copy per cell to as many as 5,300 copies per cell (5). The number of different transcripts observed in each cell population varied from 14,247 to 20,471. The bulk of the mRNA mass (75%) consisted of transcripts expressed at more than five copies per cell on average (Table 1B). In contrast, the majority (86%) of transcripts were expressed at less than 5 copies per cell, but in aggregate this low abundance class represented only 25% of the mRNA mass. This distribution was consistently observed among the different samples analyzed and was consistent with previous studies of RNA abundance classes based on RNA-DNA reassociation kinetics (Rot curves). Monte Carlo simulations revealed that our analyses had a 92% probability of detecting a transcript expressed at an average of three copies per cell (7).

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Table 1 - Summary of SAGE Analysis

A. Overall Summary

	Normal	Colon	Colon	Pancreatic	Pancreatic	
	Colon	Tumors	Cell Lines	Tumors	Cell Lines	Total
Total Tags	62,168	878,09	60,373	61,592	58,695	303,706
Unique Genes¹ GenBank²	14,721 8,753 (59)	19,690 10,490 (53)	17,092 10,193 (60)	20,471 11,547 (56)	14,247 8,922 (63)	48,741 26,339 (54)

¹ Indicates the number of different genes represented by the total tags analyzed (4).

² Indicates the number of genes that matched an entry in GenBank. The number in parentheses indicates the corresponding percentage of total unique tags.

Table 1 - Summary of SAGE Analysis

B. Summarized by Abundance Classes*

	Normal	Colon	Colon	Pancreatic	Pancreatic Cell	lla
Copies/Cell	Colon	Tumors	Cell Lines	Tumors	Lines	Total
> 500						(1)
Unique Genes	62 (29)	54 (25)	24 (19)	32 (11)	(97) 0/	(41) cc
GenBank	(56) 65	52 (96)	53 (98)	32 (100)	70 (100)	54 (98)
> 50 and < 500						
Unique Genes	645 (28)	470 (21)	618 (27)	657 (29)	585 (27)	595 (26)
GenBank	545 (84)	429 (91)	579 (94)	(60) (09)	529 (90)	553 (93)
> 5 and ≤ 50						
Unique Genes	4,569 (27)	5,011 (29)	5,733 (34)	6,146 (36)	4,895 (31)	6,209 (30)
GenBank	2,893 (63)	3,204 (64)	3,682 (64)	4,054 (66)	3,168 (65)	4,241 (68)

41,882 (25)	21,491 (51)
8,697 (16)	5,155 (59)
13,636 (24)	6,852 (50)
10,687 (20)	5,879 (55)
14,155 (25)	6,805 (48)
9,445 (16)	5,256 (56)
≥ 3 Unique Genes	GenBank

*For unique genes, the first number denotes the number of different genes (4) represented in the indicated abundance class. The number in parentheses indicates the mass fraction (X100) of total transcripts represented by the indicated abundance class. For GenBank entries, the first number indicates the number of different genes that matched an entry in GenBank in the indicated abundance class. The number in parentheses indicates the corresponding percentage of total genes.

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Many of the SAGE tags appeared to represent previously undescribed transcripts, as only 54% of the tags matched entries in GenBank (Table 1). Twenty percent of these matching transcripts corresponded to characterized mRNA sequence entries in GenBank, whereas 80% matched uncharacterized EST entries. As expected, the likelihood of a tag being present in the databases was related to abundance; GenBank matches were identified for 98% of the transcripts expressed at more than 500 copies per cell but for only 51% of the transcripts expressed at \leq 5 copies per cell. Because the SAGE data provide a quantitative assay of transcript abundance, unaffected by differences in cloning or PCR efficiency, these data provide an independent and relatively unbiased estimate of the current completeness of publicly available EST databases.

EXAMPLE 2

This example demonstrates a comparison of the expression pattern of normal colon epithelium and primary colon cancers.

Comparison of expression patterns between normal colon epithelium and primary colon cancers revealed that the majority of transcripts were expressed at similar levels (Fig. 1A). However, the expression profiles also revealed 289 transcripts that were expressed at significantly different levels [P < 0.01, (8)]. Of these 289, 181 were decreased in colon tumors compared to normal colon (average decrease 10-fold; Fig. 1B; examples in Fig. 2A). Conversely, 108 transcripts were expressed at higher levels in the colon cancers than in normal colon (average increase 13-fold; Fig. 1C; examples in Fig. 2A). Monte Carlo simulations indicated that the analysis would have detected over 95% of those transcripts expressed at a 6-fold or greater level in normal vs. tumor cells or vice versa (9). Because relatively stringent criteria were used for defining differences [P < 0.01, (8)], the number of differences reported above is likely to be an underestimate.

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EXAMPLE 3

This example demonstrates the similarities and differences between cancer cell line transcription and transcription of primary cancer tissues. To determine how many of the 289 differences were independent of the cellular microenvironment of cancers in vivo, SAGE data from CR cancer cell lines was compared to that from primary CR cancer tissues (Fig. 1B, 1C). Perhaps surprisingly, the majority of transcripts (130 of 181) that were expressed at reduced levels in cancer cells in vivo were also expressed at significantly lower levels in the cell lines (Fig. 1B). Likewise, a significant fraction of the transcripts expressed at increased levels in primary cancers were also expressed at higher levels in the CR cancer cell lines (Fig. 1C). Thus, many of the gene expression differences that distinguish normal from tumor cells in vivo persist during in vitro growth. However, despite these similarities there were also many differences. For example, only 47 of 228 genes expressed at higher levels in CR cancer cell lines were also expressed at high levels in the primary CR cancers.

In combination, comparing the expression pattern of CR cancer cells (in vivo or in vitro) to normal colon revealed 548 differentially expressed transcripts (Fig. 1B,C, Tables 2 and 3). The average difference in expression for these transcripts was 15 fold. Although the ability to detect differences is influenced by the magnitude of the variance with the power to detect smaller differences being less, 92 transcripts that were less than three fold different were identified among the 548 transcripts. However, those genes exhibiting the greatest differences in expression are likely to be the most biologically important.

EXAMPLE 4

This example demonstrates the similarities and differences between colorectal cancer transcription and pancreatic cancer transcription.

To determine whether the changes noted in CR cancers were neoplasia or cell type specific, we performed SAGE on mRNA derived from pancreatic cancers. A total of 404 transcripts were expressed at higher levels in pancreatic cancers compared to normal colon epithelium (examples in Fig. 2B). The majority (268) of these transcripts were pancreas-specific (10) (Example in Fig. 2C) although 136 were also expressed at high levels in CR cancers. These 136 transcripts constituted 47% of the 289 transcripts increased in CR cancers relative to normal colon and are likely to be related to the neoplastic process rather than to the specific cell type of origin.

EXAMPLE 5

This example demonstrates the reproducibility of the transcription patterns observed among a larger number of cancer samples.

One question that arose from these data is the potential heterogeneity of expression between individual tumors. The SAGE data were acquired from two examples of each tissue type (normal colon, primary CR cancer, CR cancer cell line, etc.). To examine the generality of these expression profiles, we arbitrarily selected 27 differentially expressed transcripts and evaluated them in six to twelve samples of normal colon and primary cancers by Northern blot analysis (11). In general, expression patterns were very reproducible among different samples. Of 10 genes with elevated expression in normal colon relative to CR cancers as determined by SAGE, each was detected in the normal colon samples and was expressed at considerably lower levels in tumors (examples in Fig. 2A). Similarly, most of the genes identified by SAGE as increased in CR or pancreatic cancers were confirmed to be reproducibly expressed in the majority of primary cancers examined by Northern blot (examples in Fig. 2A). It is important to note, however, that there were differences among the cancers, with a few cancers exhibiting particularly high or low levels of individual transcripts. Such differences in gene expression

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undoubtedly contribute to the observed heterogeneity in biological properties of cancers derived from the same organ.

EXAMPLE 6

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This example demonstrates the identities of some of the transcripts which were found to be differentially expressed in tumor and normal tissues. What are the identities of the differentially expressed genes? Of the 548 differentially expressed transcripts, 337 were tentatively identified through database comparisons. When tested, the great majority (93%) of these identifications proved to be legitimate (13), as expected from previous SAGE analyses. Although a large number of differentially expressed genes were identified, some simple patterns did emerge. For example, genes that were expressed at higher levels in normal colon epithelium than in CR tumors were often differentiation-related. These genes included liver fatty acid binding protein, cytokeratin 20, carbonic anhydrase, guanylin and uroguanylin, which are known to be important for the normal physiology or architecture of the colon epithelium (Table 2). On the other hand, genes that were increased in CR cancers were often related to the robust growth characteristics that these cells exhibit. For example, gene products associated with protein synthesis, including 48 ribosomal proteins, five elongation factors, and five genes involved in glycolysis were observed to be elevated in both CR and pancreatic cancers compared to normal colon cells. Although the majority of the transcripts could not have been predicted to be differentially expressed in cancers, several have previously been shown to be dysregulated in neoplastic The latter included IGFII, B23 nucleophosmin, the Pi form of glutathione S-transferase, and several ribosomal proteins which were all increased in cancer cells as previously reported. Likewise, Dra and gelsolin were both decreased in cancer as previously reported. Surprisingly, two widely studied oncogenes, c-fos and c-erbb3, were expressed at much higher levels in normal colon epithelium than CR cancers, in contrast to their up-regulation in transformed cells.

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In summary, these data provide basic information necessary for understanding the gene expression differences that underlie cancer phenotypes. They additionally provide a necessary framework for interpreting the significance of individual differentially expressed genes. Although this study demonstrated that a large number of such differences exist (approximately 500 at the depth of analysis employed), it was equally remarkable that the fraction of transcripts exhibiting significant differences was relatively small, representing 1.5 % of the transcripts detected in any given cell type (26). The fact that many, but not all, of the differences were preserved during in vitro culture demonstrates the utility of cultured lines for examination of some aspects of gene expression, but also provides a note of caution in relying on such lines to perfectly mimic tumors in their natural environment. Finally, the finding that hundreds of specific genes are expressed at different levels in CR cancers, and that some of these are also expressed differentially in pancreatic cancers, provides a wealth of new reagents for future biologic and diagnostic experimentation.

REFERENCES AND NOTES

- M. D. Adams, et al., Nature 377, supp. 28, 3 (1995); M. Schena, D. Shalon, R. W. Davis, P. O. Brown, Science 270, 467 (1995); J. Derisi, et al., Nature Genetics 14, 457 (1996); T. M. Gress, et al., Oncogene 13, 1819 (1996); D. J. Lockhart, et al., Nature Biotechnology 14, 1675 (1996); M. Schena, et al., Proc Natl Acad Sci USA 93, 10614 (1996).
- V. E. Velculescu, L. Zhang, B. Vogelstein, K. W. Kinzler, Science 270, 484 (1995); V. E. Velculescu, et al., Cell 88, 243 (1997).
- 3. To minimize individual variation, approximately equal numbers of tags (30,000) were derived from two different patients for each tissue. For primary tumors (two CR carcinomas and two pancreatic adenocarcinomas), RNA was isolated from portions of tumors judged to contain 60%-90% tumor cells by histopathology. The cells grown in vitro were derived from CR (SW837, Caco2) and pancreatic (ASPC-1, PL45) cancer cell lines. CR epithelial cells were isolated from sections of normal colon mucosa from two patients using EDTA as previously described [S. Nakamura, I. Kino, S. Baba, Gut 34, 1240 (1993)]. Histopathology confirmed that the isolated cells were greater than 90% epithelial. Isolation of Poly-A RNA and SAGE was performed as previously described (2). SAGE data was analyzed by means of SAGE software and GenBank Release 95 as previously described (2).
- 4. A total of 69,393 different SAGE tags were identified among the 303,706 tags analyzed. A small fraction of these different tags were likely due to sequencing errors. SAGE analysis of yeast (2), wherein the entire genomic sequence is known, demonstrated a sequencing error rate of ~ 0.7%, translating to a SAGE tag error rate of 6.8% (1 0.993¹⁰). Because these sequencing mistakes are essentially random, they do not substantially affect the analysis although they could artificially inflate the number of unique genes identified. Therefore, to be conservative, we reduced our estimate of unique genes identified by this maximum tag error rate (e.g., 6.8% of 303,706 total tags). The number of different tags derived from the same gene due to alternative splicing was assumed to be negligible.

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- 5. Abundances can be simply determined by dividing the observed number of tags for a given transcript by the total number of tags obtained. An estimate of approximately 300,000 transcripts per cell was used to convert the abundances to copies per cell [N. D. Hastie, J. O. Bishop, *Cell* 9, 761 (1976)].
- J. O. Bishop, J. G. Morton, M. Rosbash, M. Richardson, *Nature* 250, 199 (1974); B. Lewin, Gene Expression Vol 2 (John Wiley and sons,
 New York 1980).
- 7. Computer simulations indicated that analysis of 300,000 tags would yield a 92 % chance of detecting a tag for a transcript whose expression was at least three copies per cell on average among the tissues examined and assuming 300,000 transcripts per cell.
- 8. To minimize the number of assumptions and to account for the large number of comparisons being made, Monte Carlo analysis was used for determining statistical significance. The null hypothesis was that the level, kind, and distribution of transcripts were the same for cancer and normal cells. For each transcript, 100,000 simulations were performed to determine the relative likelihood due to chance alone ("p-chance") of obtaining a difference in expression equal to or greater than the observed difference, given the null hypothesis. This likelihood was converted to an absolute probability value by simulating 40 experiments in which a representative number of transcripts (27,993 transcripts in each experiment) was identified and compared. The distribution of transcripts used for these simulations was derived from the average level of expression observed in the original samples. The distribution of the p-chance scores obtained in the 40 simulated experiments (false positives) was then compared to those obtained experimentally. Based on this comparison, a maximum value of 0.0005 was chosen for p-chance. This yielded a false positive rate that was no higher than 0.01 for the least significant p-chance value below the cutoff.
- 9. Two hundred simulations assuming an abundance of 0.0001 in one sample and 0.0006 in a second sample revealed a significant difference (P < 0.01, [8]) 95% of the time.

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CLAIMS

1. A method of diagnosing colon cancer in a sample suspected of being neoplastic, comprising the steps of:

comparing the level of at least one transcript in a first sample of a tissue to a second sample, wherein the first sample is of a colonic tissue suspected of being neoplastic and the second sample is of a normal human colonic tissue, and wherein the transcript is identified by a tag selected from the group consisting of those shown in Table 3;

identifying the first sample as neoplastic when the level of the at least one transcript is found to belower in the first sample than in the second sample.

2. A method of diagnosing colon cancer in a sample suspected of being neoplastic, comprising the steps of:

comparing the level of at least one transcript in a first sample of a tissue to a second sample, wherein the first sample is of a colonic tissue suspected of being neoplastic and the second sample is of a normal human colonic tissue, and wherein the transcript is identified by a tag selected from the group consisting of those shown in Table 2;

identifying the first sample as neoplastic when the level of the at least one transcript is found to be higher in the first sample than in the second sample.

- 3. The method of claim 1 wherein a comparison of at least two of said transcripts is performed.
- 4. The method of claim 2 wherein a comparison of at least two of said transcripts is performed.

- 5. The method of claim 1 wherein a comparison of at least five of said transcripts is performed.
- 6. The method of claim 2 wherein a comparison of at least five of said transcripts is performed.
- The method of claim 1 wherein a comparison of at least ten of said transcripts is performed.
 - 8. The method of claim 2 wherein a comparison of at least ten of said transcripts is performed.
 - 9. The method of claim 1 wherein a comparison of at least twenty of said transcripts is performed.
 - 10. The method of claim 2 wherein a comparison of at least twenty of said transcripts is performed.
 - 11. The method of claim 1 wherein a comparison of at least thirty of said transcripts is performed.
- 15 12. The method of claim 2 wherein a comparison of at least thirty of said transcripts is performed.
 - 13. An isolated and purified human nucleic acid molecule which comprises a SAGE tag selected from SEQ ID NO:1-732.
 - 14. The nucleic acid molecule of claim 13 which is a cDNA molecule.

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- 15. The nucleic acid molecule of claim 13 wherein the SAGE tag is located at the 3' end of the molecule, adjacent to the 3'-most NlaIII restriction enzyme site.
- 16. An isolated nucleotide probe comprising at least 10 nucleotides of a human nucleic acid molecule, wherein the human nucleic acid molecule comprises a SAGE tag selected from SEQ ID NO: 1-732.
 - 17. The probe of claim 16 which comprises the selected SAGE tag.
 - 18. A diagnostic reagent for evaluating neoplasia of a colorectal tissue, comprising at least 2 probes according to claim 16.
- 19. The diagnostic reagent of claim 18 which comprises at least 5 probes according to claim 16.
 - 20. The diagnostic reagent of claim 18 which comprises at least 10 probes according to claim 16.
 - 21. The diagnostic reagent of claim 18 which comprises at least 20 probes according to claim 16.
 - 22. The diagnostic reagent of claim 18 which comprises at least 30 probes according to claim 16.
 - 23. A diagnostic reagent for evaluating neoplasia of a colorectal tissue, comprising at least 2 probes according to claim 17.
 - 24. A method of diagnosing pancreatic cancer in a sample suspected of being neoplastic, comprising the steps of:

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comparing the level of at least one transcript in a first sample of a tissue to a second sample, wherein the first sample is of a pancreatic tissue suspected of being neoplastic and the second sample is of a normal human colon tissue, wherein said transcript is identified by a tag selected from the group consisting of those shown Table 4;

identifying the first sample as neoplastic when the level of the at least one transcript is found to be higher in the first sample than in the second sample.

25. A method of diagnosing cancer in a sample suspected of being neoplastic, comprising the steps of:

comparing the level of at least one transcript in a first sample of a tissue to a second sample, wherein the first sample is of a tissue suspected of being neoplastic and the second sample is of a normal human tissue of the same tissue type, wherein said transcript is identified by a tag selected from the group consisting of those shown Table 5;

identifying the first sample as neoplastic when the level of the at least one transcript is found to be higher in the first sample than in the second sample.

26. A method to aid in the determination of a prognosis for a colon cancer patient, comprising the steps of:

comparing the level of at least one transcript in a first sample of a tissue to a second sample, wherein the first sample is of a neoplastic colonic tissue and the second sample is of a normal human colonic tissue, and wherein the transcript is identified by a tag selected from the group consisting of those shown in Table 3;

determining a poorer prognosis if the level of the at least one transcript is found to be lower in the first sample than in the second sample.

27. A method to aid in determining a prognosis for a patient with colon cancer, comprising the steps of:

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comparing the level of at least one transcript in a first tissue sample to a second sample, wherein the first sample is of a colonic cancer tissue and the second sample is of a normal human colonic tissue, and wherein the transcript is identified by a tag selected from the group consisting of those shown in Table 2;

determining a poorer prognosis if the level of the at least one transcript is found to be higher in the first sample than in the second sample.

28. A method of diagnosing colon cancer in a sample suspected of being neoplastic, comprising the steps of:

comparing the level of expression of at least one protein in a first sample of a tissue to a second sample, wherein the first sample is of a colonic tissue suspected of being neoplastic and the second sample is of a normal human colonic tissue, and wherein the protein is encoded by a transcript identified by a tag selected from the group consisting of those shown in Table 3;

identifying the first sample as neoplastic when the level of expression of the protein is found to be lower in the first sample than in the second sample.

29. A method of diagnosing colon cancer in a sample suspected of being neoplastic, comprising the steps of:

comparing the level of expression of at least one protein in a first sample of a tissue to a second sample, wherein the first sample is of a colonic tissue suspected of being neoplastic and the second sample is of a normal human colonic tissue, and wherein the protein is encoded by a transcript identified by a tag selected from the group consisting of those shown in Table 2;

identifying the first sample as neoplastic when expression of the protein is found to be higher in the first sample than in the second sample.

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30. A method to aid in determining a prognosis of a patient having pancreatic cancer, comprising the steps of:

comparing the level of at least one transcript in a first sample of a tissue to a second sample, wherein the first sample is of a neoplastic pancreatic tissue and the second sample is of a normal human colon tissue, wherein said transcript is identified by a tag selected from the group consisting of those shown Table 4;

determining a poorer prognosis if the level of the at least one transcript is found to be higher in the first sample than in the second sample.

31. A method to aid in providing a prognosis for a cancer patient, comprising the steps of:

comparing the level of at least one transcript in a first sample of a tissue to a second sample, wherein the first sample is of a neoplastic tissue and the second sample is of a normal human tissue of the same tissue type, wherein said transcript is identified by a tag selected from the group consisting of those shown Table 5;

determining a poorer prognosis if the level of the at least one transcript is found to be higher in the first sample than in the second sample.

32. A method of diagnosing pancreatic cancer in a sample suspected of being neoplastic, comprising the steps of:

comparing the level of expression of at least one protein encoded by a transcript in a first sample of a tissue to a second sample, wherein the first sample is of a pancreatic tissue suspected of being neoplastic and the second sample is of a normal human colon tissue, wherein said protein is encoded by a transcript identified by a tag selected from the group consisting of those shown Table 4;

identifying the first sample as neoplastic when expression of the protein is found to be higher in the first sample than in the second sample.

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33. A method of diagnosing cancer in a sample suspected of being neoplastic, comprising the steps of:

comparing the level of expression of at least one protein in a first sample of a tissue to a second sample, wherein the first sample is of a tissue suspected of being neoplastic and the second sample is of a normal human tissue, wherein said protein is encoded by a transcript identified by a tag selected from the group consisting of those shown Table 5;

identifying the first sample as neoplastic when expression of the protein is found to be higher in the first sample than in the second sample.

34. A method to aid in the determination of a prognosis for a colon cancer patient, comprising the steps of:

comparing the level of expression of at least one protein in a first sample of a tissue to a second sample, wherein the first sample is of a neoplastic colonic tissue and the second sample is of a normal human colonic tissue, and wherein the protein is encoded by a transcript identified by a tag selected from the group consisting of those shown in Table 3;

determining a poorer prognosis if the level of expression is found to be lower in the first sample than in the second sample.

35. A method to aid in determining a prognosis for a patient with colon cancer, comprising the steps of:

comparing the level of expression of at least one protein in a first tissue sample to a second sample, wherein the first sample is of a colonic cancer tissue and the second sample is of a normal human colonic tissue, and wherein the protein is encoded by a transcript identified by a tag selected from the group consisting of those shown in Table 2;

determining a poorer prognosis if the level of expression is found to be higher in the first sample than in the second sample.

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36. A method to aid in determining a prognosis of a patient having pancreatic cancer, comprising the steps of:

comparing the level of expression of at least one protein in a first sample of a tissue to a second sample, wherein the first sample is of a neoplastic pancreatic tissue and the second sample is of a normal human colon tissue, wherein said protein is encoded by a transcript identified by a tag selected from the group consisting of those shown Table 4;

determining a poorer prognosis if the level of expression is found to be higher in the first sample than in the second sample.

37. A method to aid in providing a prognosis for a cancer patient, comprising the steps of:

comparing the level of expression of at least one protein in a first sample of a tissue to a second sample, wherein the first sample is of a neoplastic tissue and the second sample is of a normal human tissue of the same tissue type, wherein said protein is encoded by a transcript identified by a tag selected from the group consisting of those shown Table 5;

determining a poorer prognosis if the level of expression is found to be higher in the first sample than in the second sample.

38. A method of treating a cancer cell, comprising the step of:

administering to a cancer cell an antibody which specifically binds to a protein encoded by a transcript identified by a tag selected from the group consisting of those shown in Tables 2, 4, and 5, wherein the antibody is linked to a cytotoxic agent.

39. An antibody linked to a cytotoxic agent, wherein the antibody specifically binds to a protein encoded by a transcript identified by a tag selected from the group consisting of those shown in Tables 2, 4, and 5.

40. A method of detecting colon cancer in a patient, comprising the steps of:

comparing the level of at least one protein in a first body sample to a second body sample, wherein the first sample is a body sample of the patient and the second sample is of a normal human, wherein the protein is encoded by a transcript identified by a tag selected from the group consisting of those shown in Table 2, wherein the first and second body sample is a sample selected from the group consisting of blood, urine, feces, sputum, and serum:

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identifying neoplasia when the level of the at least one protein is found to be higher in the first sample than in the second sample.

41. A method of detecting pancreatic cancer in a patient, comprising the steps of:

comparing the level of at least one protein encoded by a transcript in a first sample of a tissue to a second sample, wherein the first sample is of the patient and the second sample is of a normal human, wherein said protein is encoded by a transcript identified by a tag selected from the group consisting of those shown Table 4, wherein the first and second sample is a sample selected from the group consisting of blood, urine, feces, sputum, and serum;

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identifying neoplasia when the level of the at least one protein is found to be higher in the first sample than in the second sample.

42. A method of detecting cancer in a patient, comprising the steps of:

comparing the level of at least one protein in a first sample to a second sample, wherein the first sample is of patient and the second sample is of a normal human, wherein said protein is encoded by a transcript identified by a tag selected from the group consisting of those shown Table 5, wherein the first and second body sample is a sample selected from the group consisting of blood, urine, feces, sputum, and serum;

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identifying neoplasia when the level of the at least one protein is found to be higher in the first sample than in the second sample.

43. A method to aid in determining a prognosis for a patient with colon cancer, comprising the steps of:

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comparing the level of at least one protein in a first sample to a second sample, wherein the first sample is of a colonic cancer patient and the second sample is of a normal human, wherein the protein is encoded by a transcript identified by a tag selected from the group consisting of those shown in Table 2, wherein the first and second sample is a sample selected from the group consisting of blood, urine, feces, sputum, and serum;

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determining a poorer prognosis if the level of the at least one protein is found to be higher in the first sample than in the second sample.

44. A method to aid in determining a prognosis of a patient having pancreatic cancer, comprising the steps of:

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comparing the level of at least one protein in a first sample to a second sample, wherein the first sample is of a pancreatic cancer patient and the second sample is of a normal human, wherein said protein is encoded by a transcript identified by a tag selected from the group consisting of those shown Table 4, wherein said first and second sample is a sample selected from the group consisting of blood, urine, feces, sputum, and serum;

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determining a poorer prognosis if the level of the at least one protein is found to be higher in the first sample than in the second sample.

45. A method to aid in providing a prognosis for a cancer patient, comprising the steps of:

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comparing the level of expression of at least one protein in a first sample to a second sample, wherein the first sample is of a cancer patient and the second sample is of a normal human, wherein said protein is encoded by a transcript identified by a tag selected from the group consisting of those

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shown Table 5, wherein the first and second sample is a sample selected from the group consisting of blood, urine, feces, sputum, and serum;

determining a poorer prognosis if the level of the at least one protein is found to be higher in the first sample than in the second sample.

46. A method of detecting colon cancer in a patient, comprising the steps of:

comparing the level of at least one transcript in a first body sample to a second body sample, wherein the first sample is a body sample of the patient and the second sample is of a normal human, wherein the transcript is identified by a tag selected from the group consisting of those shown in Table 2, wherein the first and second body sample is a sample selected from the group consisting of blood, urine, feces, sputum, and serum;

identifying neoplasia when the level of the at least one transcript is found to be higher in the first sample than in the second sample.

47. A method of detecting pancreatic cancer in a patient, comprising the steps of:

comparing the level of at least one transcript in a first sample of a tissue to a second sample, wherein the first sample is of the patient and the second sample is of a normal human, wherein said transcript is identified by a tag selected from the group consisting of those shown Table 4, wherein the first and second sample is a sample selected from the group consisting of blood, urine, feces, sputum, and serum;

identifying neoplasia when the level of the at least one transcript is found to be higher in the first sample than in the second sample.

25 48. A method of detecting cancer in a patient, comprising the steps of:

comparing the level of at least one transcript in a first sample to
a second sample, wherein the first sample is of patient and the second sample
is of a normal human, wherein said transcript is identified by a tag selected

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from the group consisting of those shown Table 5, wherein the first and second body sample is a sample selected from the group consisting of blood, urine, feces, sputum, and serum;

identifying neoplasia when the level of the at least one transcript is found to be higher in the first sample than in the second sample.

49. A method to aid in determining a prognosis for a patient with colon cancer, comprising the steps of:

comparing the level of at least one transcript in a first sample to a second sample, wherein the first sample is of a colonic cancer patient and the second sample is of a normal human, wherein the transcript is identified by a tag selected from the group consisting of those shown in Table 2, wherein the first and second sample is a sample selected from the group consisting of blood, urine, feces, sputum, and serum;

determining a poorer prognosis if the level of the at least one transcript is found to be higher in the first sample than in the second sample.

50. A method to aid in determining a prognosis of a patient having pancreatic cancer, comprising the steps of:

comparing the level of at least one transcript in a first sample to a second sample, wherein the first sample is of a pancreatic cancer patient and the second sample is of a normal human, wherein said transcript is identified by a tag selected from the group consisting of those shown Table 4, wherein said first and second sample is a sample selected from the group consisting of blood, urine, feces, sputum, and serum;

determining a poorer prognosis if the level of the at least one transcript is found to be higher in the first sample than in the second sample.

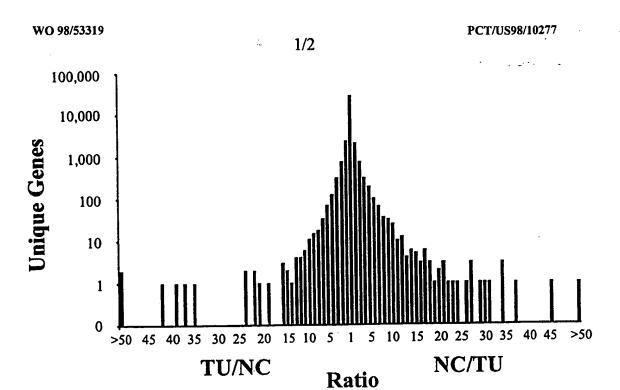
51. A method to aid in providing a prognosis for a cancer patient, comprising the steps of:

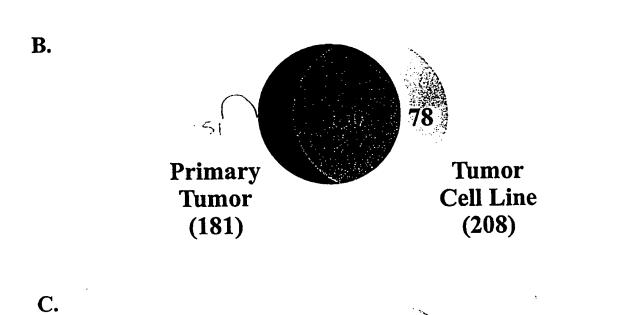
10

comparing the level of expression of at least one transcript in a first sample to a second sample, wherein the first sample is of a cancer patient and the second sample is of a normal human, wherein said transcript is identified by a tag selected from the group consisting of those shown Table 5, wherein the first and second sample is a sample selected from the group consisting of blood, urine, feces, sputum, and serum;

determining a poorer prognosis if the level of the at least one transcript is found to be higher in the first sample than in the second sample.

52. A method for screening for candidate agents that modulate the expression of a polynuleotide selected from the group consisting of the polynucleotides in SEQ ID NOS:1-732 or their respective complements, comprising contacting a test agent with a colon or pancreatic cell and monitoring expression of the polynucleotide, wherein the test agent which modifies the expression of the polynucleotide is a candidate agent.





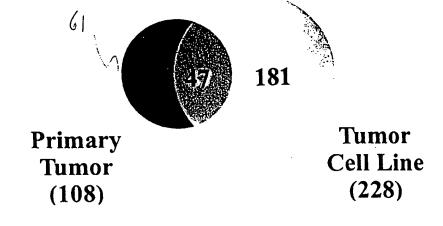


FIG. 2

A.

	1	2	SAGE Data		
	TN	r N 7	r N	T	N
			4		
H204104	•	•		11	102
H259108	•			1	37
H1000193	041)w(—	56	12
H998030	W 0		*	55	. 7

B.

	Pancreatic Tumors									mal Ion	SAGE Data	
	1	2	3	3 4 5		5 6		8	1	2	Pancreatic Tumors	Normal Colon
			-					-	-	H	Idiliois	
	-	أسا	سا	لسيا	H	7	A	<u>ل</u> ب	1	+ 1		
H294155	-	•	-	114	•	•	-)		47	0
H560056)		32	0

C.

	CR Tumors		Pancreatic Tumors			Normal Colon			SAGE Data			
	1	2	3	1	2	3	1	2	3		Pancreatic	
		—								Tumors	Tumors	Colon
H802810			-)						27	0	1
H85882				•					•	10	26	0
H618841				•		-	,	,		8	62	0